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<p>(21) International Application Number: PCT/US92/08749 (22) International Filing Date: 13 October 1992 (13.10.92) (30) Priority data: 07/776,491 11 October 1991 (11.10.91) US 07/883,944 15 May 1992 (15.05.92) US 07/953,272 29 September 1992 (29.09.92) US (71) Applicant: THE DU PONT MERCK PHARMACEUTICAL COMPANY [US/US]; 1007 Market Street, Wilmington, DE 19898 (US). (72) Inventors: LAM, Patrick, Yuk-Sun ; 6 Ridgeway Drive, Chadds Ford, PA 19317 (US). EYERMANN, Charles, Joseph ; 703 Bellevue Road, Wilmington, DE 19809 (US). HODGE, Carl, Nicholas ; 407 Lee Terrace, Wilmington, DE 19803 (US). JADHAV, Prabhakar, Kondaji ; 11 Morgan Lane, Wilmington, DE 19808 (US). DeLUCCA, George, Vincent ; 2703 Marlyn Drive, Wilmington, DE 19810 (US).</p>		<p>(74) Agents: FERGUSON, Blair, Q. et al.; The du Pont Merck Pharmaceutical Company, Legal/Patent Records Center, 1007 Market Street, Wilmington, DE 19898 (US). (81) Designated States: AU, BB, BG, BR, CA, CS, FI, HU, JP, KP, KR, LK, MG, MN, MW, NO, PL, RO, RU, SD, European patent (AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, SN, TD, TG). Published <i>With international search report.</i> <i>With amended claims.</i></p>
<p>(54) Title: CYCLIC UREAS AND ANALOGUES USEFUL AS RETROVIRAL PROTEASE INHIBITORS</p>		
<p>(57) Abstract</p> <p>This invention relates to substituted cyclic carbonyls and derivatives thereof useful as retroviral protease-inhibitors, to pharmaceutical compositions comprising such compounds, and to methods of using these compounds for treating viral infection.</p>		

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TITLECYCLIC UREAS AND ANALOGUES USEFUL AS
RETROVIRAL PROTEASE INHIBITORS

5

Cross-reference to Earlier Filed Application

This application is a continuation-in-part of U.S. Patent Application Serial Number 07/883,944, filed May 15, 1992, which is a continuation-in-part of U.S. Patent Application Serial Number 07/776,491, filed October 11, 1991.

FIELD OF THE INVENTION

15

This invention relates to substituted cyclic carbonyls and derivatives thereof useful as retroviral protease inhibitors, to pharmaceutical compositions comprising such compounds, and to methods of using these compounds for treating viral infection.

BACKGROUND OF THE INVENTION

Current treatments for viral diseases usually involve administration of compounds that inhibit viral DNA synthesis. Current treatments for AIDS (Dagani, Chem. Eng. News, November 23, 1987 pp. 41-49) involve administration of compounds such as 2',3'-dideoxycytidine, trisodium phosphonoformate, ammonium 21-tungsto-9-antimoniate, 1-b-D-ribofuranoxyl-1,2,4-triazole-3-carboxamide, 3'-azido-3'-deoxythymidine (AZT), and adriamycin that inhibit viral DNA synthesis; compounds such as AL-721 and polymannoacetate which may prevent HIV from penetrating the host cell; and compounds which treat the opportunistic infections caused by the immunosuppression resulting from HIV

infection. None of the current AIDS treatments have proven to be totally effective in treating and/or reversing the disease. In addition, many of the compounds currently used to treat AIDS cause adverse side effects including low platelet count, renal toxicity, and bone marrow cytopenia.

Proteases are enzymes which cleave proteins at specific peptide bonds. Many biological functions are controlled or mediated by proteases and their complementary protease inhibitors. For example, the protease renin cleaves the peptide angiotensinogen to produce the peptide angiotensin I. Angiotensin I is further cleaved by the protease angiotensin converting enzyme (ACE) to form the hypotensive peptide angiotensin II. Inhibitors of renin and ACE are known to reduce high blood pressure *in vivo*. However, no therapeutically useful renin protease inhibitors have been developed, due to problems of oral availability and *in vivo* stability.

The genomes of retroviruses encode a protease that is responsible for the proteolytic processing of one or more polyprotein precursors such as the pol and gag gene products. See Wellink, Arch. Virol. 98:1 (1988). Retroviral proteases most commonly process the gag precursor into the core proteins, and also process the pol precursor into reverse transcriptase and retroviral protease.

The correct processing of the precursor polyproteins by the retroviral protease is necessary for the assembly of the infectious virions. It has been shown that *in vitro* mutagenesis that produces protease-defective virus leads to the production of immature core forms which lack infectivity. See Crawford, J. Virol. 53:899 (1985); Kato et al., Virology 145:280 (1985). Therefore, retroviral protease inhibition provides an

attractive target for antiviral therapy. See Mitsuya, *Nature* 325 775 (1987).

Moore, *Biochem. Biophys. Res. Commun.*, 159 420 (1989) discloses peptidyl inhibitors of HIV protease.

- 5 Erickson, European Patent Application No. WO 89/10752 discloses derivatives of peptides which are inhibitors of HIV protease.

- 10 U.S. Patent No. 4,652,552 discloses methyl ketone derivatives of tetrapeptides as inhibitors of viral proteases. U.S. Patent No. 4,644,055 discloses halomethyl derivatives of peptides as inhibitors of viral proteases. European Patent Application No. WO 87/07836 discloses L-glutamic acid gamma-monohydroxamate as an antiviral agent.

- 15 The ability to inhibit a viral protease provides a method for blocking viral replication and therefore a treatment for viral diseases, such as AIDS, that may have fewer side effects, be more efficacious, and be less prone to drug resistance when compared to current
20 treatments.

- The topic of the present invention is substituted cyclic carbonyls and derivatives thereof, which compounds are capable of inhibiting viral protease and which compounds are believed to serve as a means of
25 combating viral diseases, such as AIDS. The substituted cyclic carbonyls and derivatives thereof of this invention provide significant improvements over protease inhibitors that are known in the art. A large number of compounds have been reported to be inhibitors of
30 proteases, such as renin, but these have suffered from lack of adequate bioavailability and are thus not useful as therapeutic agents, particularly if oral administration is desired. This poor activity has been ascribed to the relatively high molecular weight of most
35 protease inhibitors, to inadequate solubility properties, and to the presence of a number of peptide

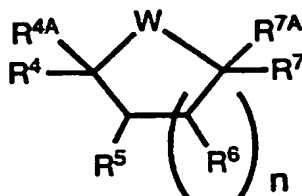
bonds, which are vulnerable to cleavage by mammalian proteases in vivo and which generally cause the molecules to be extensively bound in human serum. The substituted cyclic carbonyls and derivatives described
5 herein have a distinct advantage in this regard, in that they do not contain peptide bonds, are of low molecular weight, and can be hydrophilic yet still inhibit the viral protease enzyme.

Additionally, known inhibitors of other non-HIV
10 proteases do not inhibit HIV protease. The structure-activity requirements of such inhibitors differ from those of HIV protease inhibitors. The substituted cyclic carbonyls and derivatives of the invention are particularly useful as inhibitors of HIV protease and
15 similar retroviral proteases.

Other HIV protease inhibitors have been reported, but to date none have been shown to be clinically effective. This lack of utility is due in part to the factors discussed above for renin inhibitors,
20 particularly low bioavailability. The compounds of the invention offer a valuable solution to this problem in that they are of low molecular weight and many, therefore, have good oral absorption properties in mammals, ranging from 1-100% absolute oral
25 availability.

DETAILED DESCRIPTION OF THE INVENTION

There is provided by this invention a compound of
30 the formula (I):



(I)

or a pharmaceutically acceptable salt or prodrug form
 5 thereof wherein:

R⁴ and R⁷ are independently selected from the following
 groups:

- 10 hydrogen;
- C₁-C₈ alkyl substituted with 0-3 R¹¹;
- C₂-C₈ alkenyl substituted with 0-3 R¹¹;
- C₂-C₈ alkynyl substituted with 0-3 R¹¹;
- C₃-C₈ cycloalkyl substituted with 0-3 R¹¹;
- 15 C₆-C₁₀ bicycloalkyl substituted with 0-3 R¹¹;
- aryl substituted with 0-3 R¹²;
- a C₆-C₁₄ carbocyclic residue substituted with 0-3 R¹²;
- a heterocyclic ring system substituted with 0-2 R¹²,
- 20 R¹², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

R^{4A} and R^{7A} are independently selected from the
 25 following groups:

- hydrogen;
- C₁-C₄ alkyl substituted with halogen or C₁-C₂ alkoxy;
- 30 benzyl substituted with halogen or C₁-C₂ alkoxy;

R⁴ and R^{4A} can alternatively join to form a 5-7 membered carbocyclic ring substituted with 0-2 R¹²;

5 R⁷ and R^{7A} can alternatively join to form a 5-7 membered carbocyclic ring substituted with 0-2 R¹²;

n is 0, 1, or 2;

10 R⁵ is selected from fluoro, difluoro, =O, C₁-C₃ alkyl or -OR²⁰;

R⁶, when n is 1, is selected from: hydrogen, =O, fluoro, difluoro, C₁-C₃ alkyl or -OR²¹;

15 R⁶, when n is 2, is independently selected from: hydrogen, =O, fluoro, difluoro, C₁-C₃ alkyl or -OR²¹;

20 R⁵ and R⁶ can alternatively join to form an epoxide ring; -OCH₂SCH₂O-; -OS(=O)O-; -OC(=O)O-; -OCH₂O-; -OC(=S)O-; -OC(=O)C(=O)O-; -OC(CH₃)₂O-; -OC(OCH₃)(CH₂CH₂CH₃)O-; or any group that, when administered to a mammalian subject, cleaves to form a free dihydroxyl;

25 R²⁰ and R²¹ are independently selected from:

30 hydrogen;
C₁-C₆ alkyl substituted with 0-3 R¹¹;
C₃-C₆ alkoxyalkyl substituted with 0-3 R¹¹;
C₁-C₆ alkylcarbonyl substituted with 0-3 R¹¹;
C₁-C₆ alkoxycarbonyl substituted with 0-3 R¹¹;
benzoyl substituted with 0-3 R¹²;
phenoxycarbonyl substituted with 0-3 R¹²;
phenylaminocarbonyl substituted with 0-3 R¹²; or

any group that, when administered to a mammalian subject, cleaves to form a free hydroxyl;

R¹¹ is selected from one or more of the following:

- 5 keto, halogen, cyano, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴,
-CO₂R¹³, -OC(=O)R¹³, -OR¹³, C₂-C₆ alkoxyalkyl,
-S(O)_mR¹³, -NHC(=NH)NHR¹³, -C(=NH)NHR¹³,
-C(=O)NR¹³R¹⁴, -NR¹⁴C(=O)R¹³, =NOR¹⁴,
10 -NR¹⁴C(=O)OR¹⁴, -OC(=O)NR¹³R¹⁴, -NR¹³C(=O)NR¹³R¹⁴,
-NR¹⁴SO₂NR¹³R¹⁴, -NR¹⁴SO₂R¹³, -SO₂NR¹³R¹⁴, C₁-C₄
alkyl, C₂-C₄ alkenyl, C₃-C₆ cycloalkyl, C₃-C₆
cycloalkylmethyl;
- 15 1-3 amino acids, linked together via amide bonds
and linked to R⁴ or R⁷ via the amine or carboxylate
terminus;
- 20 a C₅-C₁₄ carbocyclic residue substituted with 0-3
R¹²;
- aryl substituted with 0-3 R¹²; or
- 25 a heterocyclic ring system substituted with 0-2
R¹², composed of 5 to 10 atoms including at least
one nitrogen, oxygen or sulfur atom;

30 R¹², when a substituent on carbon, is selected from one
or more of the following:

- phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
halogen, hydroxy, nitro, cyano, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, C₇-C₁₀ arylalkyl,
35 C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide,
oxime, boronic acid, sulfonamide, formyl, C₃-C₆

5 cycloalkoxy, $-OR^{13}$, C_1-C_4 alkyl substituted with
 $-NR^{13}R^{14}$, $-NR^{13}R^{14}$, C_2-C_6 alkoxyalkyl, C_1-C_4
hydroxyalkyl, methylenedioxy, ethylenedioxy, C_1-C_4
haloalkyl, C_1-C_4 haloalkoxy, C_1-C_4 alkoxycarbonyl,
 C_1-C_4 alkylcarbonyloxy, C_1-C_4 alkylcarbonyl, C_1-C_4
alkylcarbonylamino, $-S(O)_mR^{13}$, $-SO_2NR^{13}R^{14}$,
10 $-NHSO_2R^{14}$, $-OCH_2CO_2H$, 2-(1-morpholino)ethoxy; or

10 a 5- or 6-membered heterocyclic ring containing from
1 to 4 heteroatoms selected from oxygen, nitrogen or
sulfur;

15 or R^{12} may be a 3- or 4- carbon chain attached to
adjacent carbons on the ring to form a fused 5- or 6-
membered ring, said 5- or 6- membered ring being
optionally substituted on the aliphatic carbons with
halogen, C_1-C_4 alkyl, C_1-C_4 alkoxy, hydroxy, or
 $-NR^{13}R^{14}$; or, when R^{12} is attached to a saturated
carbon atom, it may be carbonyl or thiocarbonyl;

20 R^{12} , when a substituent on nitrogen, is selected from
one or more of the following:

25 phenyl, benzyl, phenethyl, hydroxy, C_1-C_4
hydroxyalkyl, C_1-C_4 alkoxy, C_1-C_4 alkyl, C_3-C_6
cycloalkyl, C_3-C_6 cycloalkylmethyl, $-CH_2NR^{13}R^{14}$,
 $-NR^{13}R^{14}$, C_2-C_6 alkoxyalkyl, C_1-C_4 haloalkyl, C_1-C_4
alkoxycarbonyl, $-CO_2H$, C_1-C_4 alkylcarbonyloxy,
 C_1-C_4 alkylcarbonyl;

30 R^{13} is H, phenyl, benzyl, C_1-C_6 alkyl, or C_3-C_6
alkoxyalkyl;

R^{14} is OH, H, C_1-C_4 alkyl, or benzyl;

35 R^{13} and R^{14} can alternatively join to form $-(CH_2)_4-$,

$-(CH_2)_5-$, $-CH_2CH_2N(R^{15})CH_2CH_2-$, or $-CH_2CH_2OCH_2CH_2-$;

R^{15} is H or CH_3 ;

5 m is 0, 1 or 2;

W is selected from:

- 10 $-N(R^{22})C(=Z)N(R^{23})-$;
 $-N(R^{22})S(=O)N(R^{23})-$;
 $-OC(=Z)O-$;
 $-N(R^{22})C(=Z)O-$;
 $-C(R^{25})(R^{26})C(=Z)C(R^{27})(R^{28})-$;
 $-N(R^{22})C(=Z)C(R^{27})(R^{28})-$;
 $-C(R^{25})(R^{26})C(=Z)O-$;
15 $-N(R^{22})C(=O)C(=O)N(R^{23})-$;
 $-C(R^{25})(R^{26})C(F_2)C(R^{27})(R^{28})-$;
 $-C(R^{25})(R^{26})N(CH_3)(O)C(R^{27})(R^{28})-$;
 $-C(R^{25})(R^{26})N(OR^{29})C(R^{27})(R^{28})-$;
20 $-C(R^{25})(R^{26})C(=Z)S-$;

wherein:

Z is O, S, or NR^{24} ;

25 R^{22} and R^{23} are independently selected from the following:

- hydrogen;
30 C_1 - C_8 alkyl substituted with 0-3 R^{31} ;
 C_2 - C_8 alkenyl substituted with 0-3 R^{31} ;
 C_2 - C_8 alkynyl substituted with 0-3 R^{31} ;
 C_3 - C_8 cycloalkyl substituted with 0-3 R^{31} ;
 C_6 - C_{10} bicycloalkyl substituted with 0-3 R^{31} ;
aryl substituted with 0-3 R^{32} ;

5 a C₆-C₁₄ carbocyclic residue substituted with 0-3 R³²;
a heterocyclic ring system substituted with 0-2 R³², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

R²⁴ is selected from: hydroxy; amino; C₁-C₄ alkyl; C₁-C₄ alkoxy; C₁-C₄ aminoalkyl; cyano; nitro; benzyloxy;

10 R²⁵ and R²⁷ are independently selected from the following:

15 hydrogen;
C₁-C₈ alkyl substituted with 0-3 R³¹;
C₂-C₈ alkenyl substituted with 0-3 R³¹;
C₂-C₈ alkynyl substituted with 0-3 R³¹;
C₃-C₈ cycloalkyl substituted with 0-3 R³¹;
C₆-C₁₀ bicycloalkyl substituted with 0-3 R³¹;
aryl substituted with 0-3 R³²;
20 a C₆-C₁₄ carbocyclic residue substituted with 0-3 R³²;
a heterocyclic ring system substituted with 0-2 R³², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

25 R²⁶ and R²⁸ are independently selected from:

30 hydrogen;
C₁-C₄ alkyl substituted with halogen or C₁-C₂ alkoxy;
benzyl substituted with halogen or C₁-C₂ alkoxy;

R²⁹ is selected from:

35 hydrogen;

C₁-C₄ alkyl substituted with halogen or C₁-C₂
alkoxy;
benzyl substituted with halogen or C₁-C₂ alkoxy;

- 5
alternatively, R²², R²⁵, or R²⁶, independently, can join
with R⁴ or R^{4A} to form a five- or six-membered fused
heterocyclic, aromatic, or alicyclic ring substituted
with 0-2 R¹²; and
- 10
alternatively, R²³, R²⁷, or R²⁸, independently, can join
with R⁷ or R^{7A} to form a five- or six-membered fused
heterocyclic, aromatic, or alicyclic ring substituted
with 0-2 R¹²; and
- 15
alternatively, W can join with R⁵ or R⁶ to form a three-
to seven-membered fused heterocyclic or carbocyclic ring
substituted with 0-2 R¹²;
- 20 R³¹ is selected from one or more of the following:
- keto, halogen, cyano, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴,
-CO₂R¹³,
-OC(=O)R¹³, -OR¹³, C₂-C₆ alkoxyalkyl, -S(O)_mR¹³,
25 -NHC(=NH)NHR¹³, -C(=NH)NHR¹³, -C(=O)NR¹³R¹⁴,
-NR¹⁴C(=O)R¹³, =NOR¹⁴, -NR¹⁴C(=O)OR¹⁴,
-OC(=O)NR¹³R¹⁴, -NR¹³C(=O)NR¹³R¹⁴, -NR¹⁴SO₂NR¹³R¹⁴,
-NR¹⁴SO₂R¹³, -SO₂NR¹³R¹⁴, C₁-C₄ alkyl, C₂-C₄
30 alkenyl, C₃-C₆ cycloalkyl, C₃-C₆ cycloalkylmethyl;
- 1-3 amino acids, linked together via amide bonds
and linked to R⁴ or R⁷ via the amine or carboxylate
terminus;
- 35 a C₅-C₁₄ carbocyclic residue substituted with 0-3
R³²;

aryl substituted with 0-3 R³²; or

5 a heterocyclic ring system substituted with 0-2 R³², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

R³², when a substituent on carbon, is selected from one or more of the following:

10 phenyl, benzyl, phenethyl, phenoxy, benzyloxy, halogen, hydroxy, nitro, cyano, C₁-C₄ alkyl, C₃-C₆ cycloalkyl, C₃-C₆ cycloalkylmethyl, C₇-C₁₀ arylalkyl, C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide, 15 oxime, boronic acid, sulfonamide, formyl, C₃-C₆ cycloalkoxy, -OR¹³, C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ hydroxyalkyl, methylenedioxy, ethylenedioxy, C₁-C₄ haloalkyl, C₁-C₄ haloalkoxy, C₁-C₄ alkoxycarbonyl, 20 C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, C₁-C₄ alkylcarbonylamino, -S(O)_mR¹³, -SO₂NR¹³R¹⁴, -NHSO₂R¹⁴, -OCH₂CO₂H, 2-(1-morpholino)ethoxy, -C(R¹⁴)=N(OR¹⁴); or

25 a 5- or 6-membered heterocyclic ring containing from 1 to 4 heteroatoms selected from oxygen, nitrogen or sulfur;

30 or R³² may be a 3- or 4- carbon chain attached to adjacent carbons on the ring to form a fused 5- or 6-membered ring, said 5- or 6- membered ring being optionally substituted on the aliphatic carbons with halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, hydroxy, or -NR¹³R¹⁴; or, when R³² is attached to a saturated 35 carbon atom, it may be carbonyl or thiocarbonyl;

R³², when a substituent on nitrogen, is selected from one or more of the following:

5 phenyl, benzyl, phenethyl, hydroxy, C₁-C₄
hydroxyalkyl, C₁-C₄ alkoxy, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, -CH₂NR¹³R¹⁴,
-NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ haloalkyl, C₁-C₄
alkoxycarbonyl, -CO₂H, C₁-C₄ alkylcarbonyloxy,
10 C₁-C₄ alkylcarbonyl, -C(R¹⁴)=N(OR¹⁴);

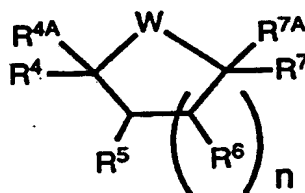
provided that:

R⁴, R^{4A}, R⁷ and R^{7A} are not all hydrogen;

15 when W is -OC(=Z)O-, R⁴ and R⁷ are not hydrogen;

when R⁴, R^{4A} are hydrogen, at least one of the
following is not hydrogen: R²², R²³, R²⁵, R²⁶,
20 R²⁷ and R²⁸.

Preferred compounds of this invention are compounds
of the formula (I):



25

(I)

or a pharmaceutically acceptable salt or prodrug form
thereof wherein:

30

R⁴ and R⁷ are independently selected from the following groups:

- 5 hydrogen;
 C₁-C₄ alkyl substituted with 0-3 R¹¹;
 C₃-C₄ alkenyl substituted with 0-3 R¹¹;
 C₃-C₄ alkynyl substituted with 0-3 R¹¹;

10 R^{4A} and R^{7A} are hydrogen;

 n is 0 or 1;

 R⁵ is selected from fluoro, difluoro, =O, or -OR²⁰;

15 R⁶ is selected from: hydrogen, =O, fluoro, difluoro, or -OR²¹;

20 R⁵ and R⁶ can alternatively join to form an epoxide ring; -OCH₂SCH₂O-; -OS(=O)O-; -OC(=O)O-; -OCH₂O-; -OC(=S)O-; -OC(=O)C(=O)O-; -OC(CH₃)₂O-; -OC(OCH₃)(CH₂CH₂CH₃)O-; or any group that, when administered to a mammalian subject, cleaves to form a free dihydroxyl;

25 R²⁰ and R²¹ are independently selected from:

- 30 hydrogen;
 C₁-C₆ alkylcarbonyl;
 C₁-C₆ alkoxy carbonyl;
 benzoyl; or
 any group that, when administered to a mammalian subject, cleaves to form a free hydroxyl;

35 R¹¹ is selected from one or more of the following:

5 keto, halogen, cyano, $-\text{CH}_2\text{NR}^{13}\text{R}^{14}$, $-\text{NR}^{13}\text{R}^{14}$,
-CO₂R¹³,
-OC(=O)R¹³, -OR¹³, C₂-C₄ alkoxyalkyl, -S(O)_mR¹³,
C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆ cycloalkyl;

a C₅-C₁₄ carbocyclic residue substituted with 0-3
R¹²;

10 aryl substituted with 0-3 R¹²; or

a heterocyclic ring system substituted with 0-2
R¹², composed of 5 to 10 atoms including at least
one nitrogen, oxygen or sulfur atom;

15

R¹², when a substituent on carbon, is selected from one
or more of the following:

20 phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
halogen, hydroxy, nitro, cyano, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, C₇-C₁₀ arylalkyl,
C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide,
oxime, boronic acid, sulfonamide, formyl, C₃-C₆
25 cycloalkoxy, -OR¹³, C₁-C₄ alkyl substituted with
-NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄
hydroxyalkyl, methylenedioxy, ethylenedioxy, C₁-C₄
haloalkyl, C₁-C₄ haloalkoxy, C₁-C₄ alkoxycarbonyl,
C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, C₁-C₄
30 alkylcarbonylamino, -S(O)_mR¹³,
-SO₂NR¹³R¹⁴, -NHSO₂R¹⁴; or

a 5- or 6-membered heterocyclic ring containing from
1 to 4 heteroatoms selected from oxygen, nitrogen or
35 sulfur;

or R¹² may be a 3- or 4- carbon chain attached to adjacent carbons on the ring to form a fused 5- or 6-membered ring, said 5- or 6- membered ring being optionally substituted on the aliphatic carbons with halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, hydroxy, or -NR¹³R¹⁴; or, when R¹² is attached to a saturated carbon atom, it may be carbonyl or thiocarbonyl;

R¹², when a substituent on nitrogen, is selected from one or more of the following:

phenyl, benzyl, phenethyl, hydroxy, C₁-C₄ hydroxyalkyl, C₁-C₄ alkoxy, C₁-C₄ alkyl, C₃-C₆ cycloalkyl, C₃-C₆ cycloalkylmethyl, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxycarbonyl, C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, -CO₂H;

R¹³ is H, C₁-C₆ alkyl, or C₃-C₆ alkoxyalkyl;

20

R¹⁴ is OH, H, C₁-C₄ alkyl, or benzyl;

R¹³ and R¹⁴ can alternatively join to form -(CH₂)₄-, -(CH₂)₅-, -CH₂CH₂N(R¹⁵)CH₂CH₂-, or -CH₂CH₂OCH₂CH₂-;

25

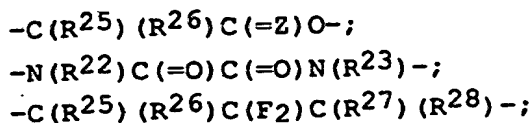
R¹⁵ is H or CH₃;

m is 0, 1 or 2;

30 W is selected from:

-N(R²²)C(=Z)N(R²³)-;
 -N(R²²)C(=Z)O-;
 -C(R²⁵)(R²⁶)C(=Z)C(R²⁷)(R²⁸)-;
 -N(R²²)C(=Z)C(R²⁷)(R²⁸)-;

35



5 wherein:

Z is O, S, N-CN, N-OH, N-OCH₃;

10 R²² and R²³ are independently selected from the following:

hydrogen;
C₁-C₈ alkyl substituted with 0-3 R³¹;
C₃-C₈ alkenyl substituted with 0-3 R³¹;
15 C₃-C₈ alkynyl substituted with 0-3 R³¹;
C₃-C₆ cycloalkyl substituted with 0-3 R³¹;

R²⁵ and R²⁷ are independently selected from the following:

20

hydrogen;
C₁-C₈ alkyl substituted with 0-3 R³¹;
C₂-C₈ alkenyl substituted with 0-3 R³¹;
C₃-C₈ alkynyl substituted with 0-3 R³¹;

25

R²⁶ and R²⁸ are hydrogen;

R³¹ is selected from one or more of the following:

30

keto, halogen, cyano, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴,
-CO₂R¹³, -OC(=O)R¹³, -OR¹³, C₂-C₄ alkoxyalkyl,
-S(O)_mR¹³, C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆
cycloalkyl;

a C₅-C₁₄ carbocyclic residue substituted with 0-3 R¹²;

aryl substituted with 0-3 R³²; or

5

a heterocyclic ring system substituted with 0-2 R³², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

10

R³², when a substituent on carbon, is selected from one or more of the following:

15 phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
halogen, hydroxy, nitro, cyano, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, C₇-C₁₀ arylalkyl,
C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide,
oxime, boronic acid, sulfonamide, formyl, C₃-C₆
20 cycloalkoxy, -OR¹³, C₁-C₄ alkyl substituted with
-NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄
hydroxyalkyl, methylenedioxy, ethylenedioxy, C₁-C₄
haloalkyl, C₁-C₄ haloalkoxy, C₁-C₄ alkoxy carbonyl,
C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, C₁-C₄
alkylcarbonylamino, -S(O)_mR¹³,
25 -SO₂NR¹³R¹⁴, -NHSO₂R¹⁴, -C(R¹⁴)=N(OR¹⁴); or

a 5- or 6-membered heterocyclic ring containing from 1 to 4 heteroatoms selected from oxygen, nitrogen or sulfur;

30

or R³² may be a 3- or 4- carbon chain attached to adjacent carbons on the ring to form a fused 5- or 6-membered ring, said 5- or 6- membered ring being optionally substituted on the aliphatic carbons with
35 halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, hydroxy, or

-NR¹³R¹⁴; or, when R³² is attached to a saturated carbon atom, it may be carbonyl or thiocarbonyl;

R³², when a substituent on nitrogen, is selected from
5 one or more of the following:

phenyl, benzyl, phenethyl, hydroxy, C₁-C₄
hydroxyalkyl, C₁-C₄ alkoxy, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, -CH₂NR¹³R¹⁴,
10 -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ haloalkyl, C₁-C₄
alkoxycarbonyl, C₁-C₄ alkylcarbonyloxy, C₁-C₄
alkylcarbonyl, -CO₂H, -C(R¹⁴)=N(OR¹⁴);

provided that:

15

R⁴, R^{4A}, R⁷, and R^{7A} are not all hydrogen;

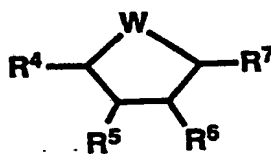
when W is -OC(=Z)O-, R⁴ and R⁷ are not hydrogen;

20

when R⁴ and R^{4A} are hydrogen, at least one of the
following is not hydrogen: R²², R²³, R²⁵, R²⁶, R²⁷
and R²⁸.

25

Further preferred compounds of the invention of
formula (I) are compounds of formula (II):



30

(II)

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R⁴ and R⁷ are independently selected from the following groups:

- 5 hydrogen;
 C₁-C₄ alkyl substituted with 0-3 R¹¹;
 C₃-C₄ alkenyl substituted with 0-3 R¹¹;

R⁵ is -OR²⁰;

- 10 R⁶ is hydrogen or -OR²¹;

R²⁰ and R²¹ are independently hydrogen or any group that, when administered to a mammalian subject, cleaves
15 to form a free hydroxyl;

R¹¹ is selected from one or more of the following:

- 20 keto, halogen, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, -OR¹³, C₂-C₄
 alkoxyalkyl, C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆
 cycloalkyl;

 aryl substituted with 0-3 R¹²; or

- 25 a heterocyclic ring system substituted with 0-2
 R¹², composed of 5 to 10 atoms including at least
 one nitrogen, oxygen or sulfur atom;

- 30 R¹², when a substituent on carbon, is selected from one
 or more of the following:

- 35 phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
 halogen, C₁-C₄ alkyl, C₇-C₁₀ arylalkyl, C₁-C₄ alkoxy,
 -CO₂H, hydroxamic acid, hydrazide, oxime, boronic
 acid, sulfonamide, formyl, C₃-C₆ cycloalkoxy, -OR¹³,

C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴, methylenedioxy, C₁-C₄ haloalkyl, C₁-C₄ alkylcarbonyl, C₁-C₄ alkylcarbonylamino, hydroxy, hydroxymethyl; or

5 a 5- or 6-membered heterocyclic ring containing from 1 to 4 heteroatoms selected from oxygen, nitrogen or sulfur;

10 R¹², when a substituent on nitrogen, is selected from benzyl or methyl;

R¹³ is H, C₁-C₂ alkyl, or C₃-C₆ alkoxyalkyl;

R¹⁴ is OH, H or C₁-C₂ alkyl;

15 R¹³ and R¹⁴ can alternatively join to form -(CH₂)₄-, -(CH₂)₅-, -CH₂CH₂N(R¹⁵)CH₂CH₂-, or -CH₂CH₂OCH₂CH₂-;

W is selected from:

20 -N(R²²)C(=Z)N(R²³)-;
-C(R²⁵)(R²⁶)C(=Z)C(R²⁷)(R²⁸)-;
-N(R²²)C(=Z)C(R²⁷)(R²⁸)-;
-C(R²⁵)(R²⁶)C(=Z)O-;

25 wherein:

Z is O, S, or N-CN;

30 R²² and R²³ are independently selected from the following:

hydrogen;
C₁-C₄ alkyl substituted with 0-3 R³¹;
35 C₃-C₄ alkenyl substituted with 0-3 R³¹;

R²⁵ and R²⁷ are independently selected from the following:

- 5 hydrogen;
 C₁-C₄ alkyl substituted with 0-3 R³¹;
 C₃-C₄ alkenyl substituted with 0-3 R³¹;

R²⁶ and R²⁸ are hydrogen;

- 10 R³¹ is selected from one or more of the following:

 keto, halogen, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, -OR¹³, C₂-C₄
 alkoxyalkyl, C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆
 cycloalkyl;

- 15 aryl substituted with 0-3 R³²; or

 a heterocyclic ring system substituted with 0-2
 R³², composed of 5 to 10 atoms including at least
20 one nitrogen, oxygen or sulfur atom;

R³², when a substituent on carbon, is selected from one or more of the following:

- 25 phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
 halogen, C₁-C₄ alkyl, C₇-C₁₀ arylalkyl, C₁-C₄ alkoxy,
 -CO₂H, hydroxamic acid, hydrazide, oxime, boronic
 acid, sulfonamide, formyl, C₃-C₆ cycloalkoxy, -OR¹³,
 C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴,
30 methylenedioxy, C₁-C₄ haloalkyl, C₁-C₄ alkylcarbonyl,
 C₁-C₄ alkylcarbonylamino, hydroxy, hydroxymethyl,
 -C(R¹⁴)=N(OR¹⁴); or

a 5- or 6-membered heterocyclic ring containing from 1 to 4 heteroatoms selected from oxygen, nitrogen or sulfur;

- 5 R³², when a substituent on nitrogen, is selected from benzyl or methyl;

provided that:

- 10 R⁴ and R⁷ are not both hydrogen;

when R⁴ is hydrogen, at least one of the following is not hydrogen: R²², R²³, R²⁵, R²⁶, R²⁷ and R²⁸.

15

More preferred compounds of the present invention are compounds of the further preferred scope above, wherein:

- 20 R⁴ and R⁷ are independently selected from the following groups:

hydrogen;

C₁-C₃ alkyl substituted with 0-1 R¹¹;

25

R⁵ is -OR²⁰;

R⁶ is hydrogen or -OR²¹;

- 30 R²⁰ and R²¹ are independently hydrogen or any group that, when administered to a mammalian subject, cleaves to form a free hydroxyl;

R¹¹ is selected from one or more of the following:

35

halogen, -OR¹³, C₁-C₄ alkyl, C₃-C₅ cycloalkyl;

5 aryl substituted with 0-2 R¹²; or
 a heterocyclic ring system chosen from pyridyl,
 pyrimidinyl, triazinyl, furanyl, thienyl, pyrrolyl,
 pyrazolyl, imidazolyl, tetrazolyl, benzofuranyl,
 indolyl, quinolinyl, isoquinolinyl;

10 R¹², when a substituent on carbon, is selected from one
 or more of the following:

benzyloxy, halogen, methyl, C₁-C₄ alkoxy, CF₃,
 2-(1-morpholino)ethoxy, -CO₂H, hydroxamic acid,
 15 hydrazide, oxime, cyano, boronic acid, sulfonamide,
 formyl, C₃-C₆ cycloalkoxy, C₁-C₄ alkyl substituted
 with -NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl; or

R¹², when a substituent on nitrogen, is methyl;

20 R¹³ is H or methyl;

R¹⁴ is OH, H or methyl;

R¹³ and R¹⁴ can alternatively join to form -(CH₂)₄-,
 25 -(CH₂)₅-, -CH₂CH₂N(R¹⁵)CH₂CH₂-, or -CH₂CH₂OCH₂CH₂-;

W is -N(R²²)C(=O)N(R²³)- or -N(R²²)C(=N-CN)N(R²³)-;

30 R²² and R²³ are independently selected from the
 following:

hydrogen;
 C₁-C₄ alkyl substituted with 0-1 R³¹;
 35 C₃-C₄ alkenyl substituted with 0-1 R³¹;

R³¹ is selected from one or more of the following:

halogen, -OR¹³, C₁-C₄ alkyl, C₃-C₅ cycloalkyl;

5 aryl substituted with 0-2 R³²; or

a heterocyclic ring system chosen from pyridyl,
pyrimidinyl, triazinyl, furanyl, thienyl, pyrrolyl,
pyrazolyl, imidazolyl, tetrazolyl, benzofuranyl,
10 indolyl, quinolinyl, isoquinolinyl;

R³², when a substituent on carbon, is selected from one
or more of the following:

15 benzyloxy, halogen, methyl, C₁-C₄ alkoxy, CF₃,
2-(1-morpholino)ethoxy, -CO₂H, hydroxamic acid,
hydrazide, oxime, cyano, boronic acid, sulfonamide,
formyl, C₃-C₆ cycloalkoxy, C₁-C₄ alkyl substituted
with -NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl,
20 -C(R¹⁴)=N(OR¹⁴); or

R³², when a substituent on nitrogen, is methyl;

provided that:

25 when R⁴ is hydrogen, R⁷ is not hydrogen;

when R⁴ is hydrogen, at least one of the following
is not hydrogen: R²² and R²³.

30 Still more preferred compounds of the present
invention are compounds of the further preferred scope
above, wherein:

35 R⁴ and R⁷ are benzyl;

R⁵ is -OH;

R⁶ is hydrogen or -OH;

5

R¹³ is H or methyl;

R¹⁴ is H or methyl;

10 W is -N(R²²)C(=O)N(R²³)- or -N(R²²)C(=N-CN)N(R²³)-;

R²² and R²³ are independently selected from the following:

15

hydrogen;

C₁-C₄ alkyl substituted with 0-1 R³¹;

R³¹ is selected from one or more of the following:

20

C₃-C₅ cycloalkyl;

aryl substituted with 0-2 R³²; or

25

a heterocyclic ring system chosen from pyridyl, thienyl, quinoliny, or isoquinoliny;

R³², when a substituent on carbon, is selected from one or more of the following:

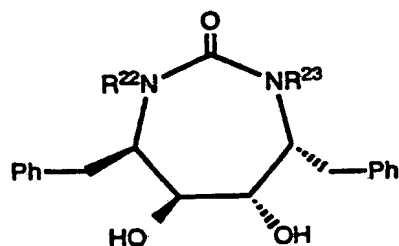
30

-CONH₂, -CO₂H, -CHO, -CH₂NHOH, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl, -C(R¹⁴)=N(OR¹⁴); or

R³², when a substituent on nitrogen, is methyl.

35

Also preferred are compounds of formula (IIa):



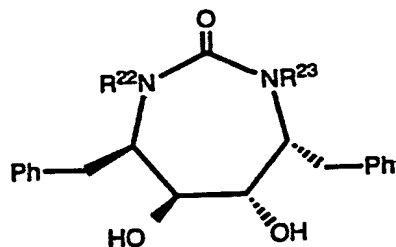
(IIa)

wherein R²² and R²³ are independently selected from the
 5 group consisting of:

hydrogen, allyl, propyl, cyclopropylmethyl,
 n-butyl, i-butyl, CH₂CH=CH(CH₃)₂, pyridylmethyl,
 methallyl, n-pentyl, i-pentyl, hexyl, benzyl,
 10 pyridylmethyl, isoprenyl, propargyl, picolinyl,
 methoxyethyl, cyclohexylmethyl, dimethyl-butyl,
 ethoxyethyl, methyl-oxazolinylmethyl,
 naphthylmethyl, methyloxazolinylmethyl,
 vinyloxyethyl, pentafluorobenzyl, quinolinylmethyl,
 15 carboxybenzyl, chloro-thienyl, picolinyl,
 benzyloxybenzyl, phenylbenzyl, adamantylethyl,
 cyclopropylmethoxybenzyl, ethoxybenzyl,
 hydroxybenzyl, hydroxymethylbenzyl, aminobenzyl,
 formylbenzyl, cyanobenzyl, cinnamyl,
 20 allyloxybenzyl, fluorobenzyl, cyclobutylmethyl,
 formaldoximebenzyl, cyclopentylmethyl, nitrobenzyl,
 nitrilobenzyl, carboxamidobenzyl,
 carbomethoxybenzyl, tetrazolylbenzyl, and
 dimethylallyl.

25

Specifically preferred are compounds of formula
 (IIa):



(IIa)

selected from the group consisting of:

5

the compound of formula (IIa) wherein R²² is allyl and R²³ is allyl;

the compound of formula (IIa) wherein R²² is propyl and R²³ is propyl;

10

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cyclopropylmethyl;

15

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is n-butyl;

the compound of formula (IIa) wherein R²² is CH₂CH=CH(CH₃)₂ and R²³ is CH₂CH=CH(CH₃)₂;

20

the compound of formula (IIa) wherein R²² is i-pentyl and R²³ is i-pentyl;

the compound of formula (IIa) wherein R²² is 4-pyridylmethyl and R²³ is 4-pyridylmethyl;

25

the compound of formula (IIa) wherein R²² is 2-methallyl and R²³ is 2-methallyl;

30

the compound of formula (IIa) wherein R²² is n-pentyl and R²³ is n-pentyl;

- the compound of formula (IIa) wherein R²² is i-butyl and R²³ is i-butyl;
- 5 the compound of formula (IIa) wherein R²² is benzyl and R²³ is benzyl;
- the compound of formula (IIa) wherein R²² is 3-pyridylmethyl and R²³ is 3-pyridylmethyl;
- 10 the compound of formula (IIa) wherein R²² is allyl and R²³ is isoprenyl;
- the compound of formula (IIa) wherein R²² is 3-propargyl and R²³ is 3-propargyl;
- 15 the compound of formula (IIa) wherein R²² is 2-picolinyl and R²³ is 2-picolinyl;
- the compound of formula (IIa) wherein R²² is 2-methoxyethyl and R²³ is 2-methoxyethyl;
- 20 the compound of formula (IIa) wherein R²² is cyclohexylmethyl and R²³ is cyclohexylmethyl;
- 25 the compound of formula (IIa) wherein R²² is 3,3-dimethyl-1-butyl and R²³ is 3,3-dimethyl-1-butyl;
- the compound of formula (IIa) wherein R²² is 2-ethoxyethyl and R²³ is 2-ethoxyethyl;
- 30 the compound of formula (IIa) wherein R²² is 3-methyl-5-oxazolinylmethyl and R²³ is hydrogen;
- the compound of formula (IIa) wherein R²² is 1-naphthylmethyl and R²³ is 1-naphthylmethyl;
- 35

the compound of formula (IIa) wherein R²² is
3-methyloxazolinylmethyl and R²³ is
3-methyloxazolinylmethyl;

5

the compound of formula (IIa) wherein R²² is
2-vinyloxyethyl and R²³ is 2-vinyloxyethyl;

10 the compound of formula (IIa) wherein R²² is
2,3,4,5,6-pentafluorobenzyl and R²³ is
2,3,4,5,6-pentafluorobenzyl;

the compound of formula (IIa) wherein R²² is
benzyl and R²³ is 2-quinolinylmethyl;

15

the compound of formula (IIa) wherein R²² is
4-carboxybenzyl and R²³ is 4-carboxybenzyl;

20 the compound of formula (IIa) wherein R²² is
5-chloro-2-thienyl and R²³ is 5-chloro-2-thienyl;

the compound of formula (IIa) wherein R²² is
2-quinolinylmethyl and R²³ is 2-quinolinylmethyl;

25

the compound of formula (IIa) wherein R²² is
2-propyl and R²³ is 2-picolinyl;

the compound of formula (IIa) wherein R²² is
3-benzyloxybenzyl and R²³ is 3-benzyloxybenzyl;

30

the compound of formula (IIa) wherein R²² is
4-phenylbenzyl and R²³ is phenylbenzyl;

35 the compound of formula (IIa) wherein R²² is
2-adamantylethyl and R²³ is 2-adamantylethyl;

the compound of formula (IIa) wherein R^{22} is hydrogen and R^{23} is cyclopropylmethyl;

5 the compound of formula (IIa) wherein R^{22} is 2-picolinyl and R^{23} is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is hydrogen;

10 the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is 2-picolinyl;

the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is 4-picolinyl;

15 the compound of formula (IIa) wherein R^{22} is 3-benzyloxybenzyl and R^{23} is 3-benzyloxybenzyl;

20 the compound of formula (IIa) wherein R^{22} is 3-cyclopropylmethoxybenzyl and R^{23} is 3-cyclopropylmethoxybenzyl;

the compound of formula (IIa) wherein R^{22} is 3-ethoxybenzyl and R^{23} is 3-ethoxybenzyl;

25 the compound of formula (IIa) wherein R^{22} is 4-benzyloxybenzyl and R^{23} is 4-benzyloxybenzyl.

30 the compound of formula (IIa) wherein R^{22} is 3-hydroxybenzyl and R^{23} is 3-hydroxybenzyl;

the compound of formula (IIa) wherein R^{22} is 4-hydroxybenzyl and R^{23} is 4-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is
3-hydroxymethylbenzyl and R²³ is
3-hydroxymethylbenzyl;

5 the compound of formula (IIa) wherein R²² is
4-hydroxymethylbenzyl and R²³ is
4-hydroxymethylbenzyl;

10 the compound of formula (IIa) wherein R²² is
3-aminobenzyl and R²³ is 3-aminobenzyl;

the compound of formula (IIa) wherein R²² is
3-carboxylbenzyl and R²³ is 3-carboxylbenzyl;

15 the compound of formula (IIa) wherein R²² is
3-formylbenzyl and R²³ is 3-formylbenzyl;

the compound of formula (IIa) wherein R²² is
3-cyanobenzyl and R²³ is 3-cyanobenzyl;

20 the compound of formula (IIa) wherein R²² is
2-naphthylmethyl and R²³ is 2-naphthylmethyl;

25 the compound of formula (IIa) wherein R²² is
n-butyl and R²³ is benzyl;

the compound of formula (IIa) wherein R²² is
allyl and R²³ is cyclopropylmethyl;

30 the compound of formula (IIa) wherein R²² is
n-butyl and R²³ is cyclopropylmethyl

the compound of formula (IIa) wherein R²² is
3-methylallyl and R²³ is benzyl;

35

the compound of formula (IIa) wherein R^{22} is benzyl and R^{23} is ethyl;

5 the compound of formula (IIa) wherein R^{22} is benzyl and R^{23} is 4-picolinyl;

the compound of formula (IIa) wherein R^{22} is cyclopropylmethyl and R^{23} is 4-picolinyl;

10 the compound of formula (IIa) wherein R^{22} is benzyl and R^{23} is cyclopentylmethyl;

the compound of formula (IIa) wherein R^{22} is cyclopropylmethyl and R^{23} is cyclopentylmethyl;

15 the compound of formula (IIa) wherein R^{22} is benzyl and R^{23} is n-propyl;

the compound of formula (IIa) wherein R^{22} is cyclopropylmethyl and R^{23} is cinnamyl;

the compound of formula (IIa) wherein R^{22} is cyclopropylmethyl and R^{23} is 2-naphthylmethyl;

25 the compound of formula (IIa) wherein R^{22} is cyclopentylmethyl and R^{23} is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is benzyl and R^{23} is 2-naphthylmethyl;

30 the compound of formula (IIa) wherein R^{22} is cyclopropylmethyl and R^{23} is 2-picolinyl;

the compound of formula (IIa) wherein R^{22} is 3-cyanobenzyl and R^{23} is 3-cyanobenzyl;

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the compound of formula (IIa) wherein R²² is 3-allyl and R²³ is 2-naphthylmethyl;

5 the compound of formula (IIa) wherein R²² is n-propyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is 2-naphthylmethyl;

10 the compound of formula (IIa) wherein R²² is H and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is 4-picolinyl and R²³ is 2-naphthylmethyl;

15 the compound of formula (IIa) wherein R²² is 3-allyl and R²³ is cyclopentylmethyl;

20 the compound of formula (IIa) wherein R²² is 3-allyl and R²³ is 2-quinolinylmethyl;

the compound of formula (IIa) wherein R²² is 3-picolinyl and R²³ is cyclopropylmethyl;

25 the compound of formula (IIa) wherein R²² is 3-picolinyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is 3-allyloxybenzyl and R²³ is 3-allyloxybenzyl;

30 the compound of formula (IIa) wherein R²² is 3-allyloxybenzyl and R²³ is 3-hydroxybenzyl;

35 the compound of formula (IIa) wherein R²² is 3-picolinyl and R²³ is 3-picolinyl;

- the compound of formula (IIa) wherein R^{22} is 2-naphthylmethyl and R^{23} is 4-fluorobenzyl;
- 5 the compound of formula (IIa) wherein R^{22} is 3-carbomethoxybenzyl and R^{23} is 3-carbomethoxybenzyl;
- the compound of formula (IIa) wherein R^{22} is 4-formylbenzyl and R^{23} is 4-formylbenzyl;
- 10 the compound of formula (IIa) wherein R^{22} is 4-cyanobenzyl and R^{23} is 4-cyanobenzyl;
- the compound of formula (IIa) wherein R^{22} is 4-hydroxybenzyl and R^{23} is n-propyl;
- 15 the compound of formula (IIa) wherein R^{22} is 3-hydroxybenzyl and R^{23} is n-propyl;
- the compound of formula (IIa) wherein R^{22} is 3-carboxybenzyl and R^{23} is 3-carboxybenzyl;
- 20 the compound of formula (IIa) wherein R^{22} is cyclobutylmethyl and R^{23} is cyclobutylmethyl;
- 25 the compound of formula (IIa) wherein R^{22} is cyclopentylmethyl and R^{23} is cyclopentylmethyl;
- the compound of formula (IIa) wherein R^{22} is n-butyl and R^{23} is 3-methylallyl;
- 30 the compound of formula (IIa) wherein R^{22} is n-butyl and R^{23} is cyclopentylmethyl;
- the compound of formula (IIa) wherein R^{22} is 3-formaldoximebenzyl and R^{23} is 3-formaldoximebenzyl;
- 35

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 3-hydroxybenzyl;

5 the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 4-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is 3-(N-methylamino)benzyl and R²³ is 3-(N-methylamino)benzyl;

10 the compound of formula (IIa) wherein R²² is 3-acetylbenzyl and R²³ is 3-acetylbenzyl;

15 the compound of formula (IIa) wherein R²² is 3-hydroxylaminobenzyl and R²³ is 3-hydroxylaminobenzyl;

20 the compound of formula (IIa) wherein R²² is 2-naphthylmethyl and R²³ is 3-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is 4-hydroxymethylbenzyl and R²³ is 3-hydroxybenzyl;

25 the compound of formula (IIa) wherein R²² is N-methyl-(3-amido)benzyl and R²³ is N-methyl-(3-amido)benzyl;

the compound of formula (IIa) wherein R²² is N-methyl-(3-amido)benzyl and R²³ is 3-(amidino)benzyl;

30 the compound of formula (IIa) wherein R²² is 3-(5-tetrazolyl)benzyl and R²³ is cyclopropylmethyl;

35 the compound of formula (IIa) wherein R²² is 3-(5-tetrazolyl)benzyl and R²³ is 3-(5-tetrazolyl)benzyl;

the compound of formula (IIa) wherein R²² is phenylmethyl-3-boronic acid and R²³ is phenylmethyl-3-boronic acid.

5

More specifically preferred are the following compounds of formula (I), which have IC₅₀ or K_i values of less than about 10 nM for inhibiting HIV protease:

10

the compound of formula (IIa) wherein R²² is allyl and R²³ is allyl;

15

the compound of formula (IIa) wherein R²² is propyl and R²³ is propyl;

20

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cyclopropylmethyl;

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is n-butyl;

25

the compound of formula (IIa) wherein R²² is CH₂CH=CH(CH₃)₂ and R²³ is CH₂CH=CH(CH₃)₂;

the compound of formula (IIa) wherein R²² is i-pentyl and R²³ is i-pentyl;

30

the compound of formula (IIa) wherein R²² is 2-methylallyl and R²³ is 2-methylallyl;

the compound of formula (IIa) wherein R²² is n-pentyl and R²³ is n-pentyl;

35

the compound of formula (IIa) wherein R²² is benzyl
and R²³ is benzyl;

5 the compound of formula (IIa) wherein R²² is allyl
and R²³ is isoprenyl;

the compound of formula (IIa) wherein R²² is
3-hydroxybenzyl and R²³ is 3-hydroxybenzyl;

10 the compound of formula (IIa) wherein R²² is
4-hydroxybenzyl and R²³ is 4-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is
3-hydroxymethylbenzyl and R²³ is
15 3-hydroxymethylbenzyl;

the compound of formula (IIa) wherein R²² is
4-hydroxymethylbenzyl and R²³ is
4-hydroxymethylbenzyl;

20 the compound of formula (IIa) wherein R²² is
3-aminobenzyl and R²³ is 3-aminobenzyl;

the compound of formula (IIa) wherein R²² is
25 3-carboxylbenzyl and R²³ is 3-carboxylbenzyl;

the compound of formula (IIa) wherein R²² is
3-formylbenzyl and R²³ is 3-formylbenzyl;

30 the compound of formula (IIa) wherein R²² is
3-cyanobenzyl and R²³ is 3-cyanobenzyl;

the compound of formula (IIa) wherein R²² is
2-naphthylmethyl and R²³ is 2-naphthylmethyl;
35

the compound of formula (IIa) wherein R^{22} is
n-butyl and R^{23} is benzyl;

5 the compound of formula (IIa) wherein R^{22} is
allyl and R^{23} is cyclopropylmethyl;

the compound of formula (IIa) wherein R^{22} is
n-butyl and R^{23} is cyclopropylmethyl

10 the compound of formula (IIa) wherein R^{22} is
3-methylallyl and R^{23} is benzyl;

the compound of formula (IIa) wherein R^{22} is
benzyl and R^{23} is ethyl;

15 the compound of formula (IIa) wherein R^{22} is
benzyl and R^{23} is 4-picolinyl;

20 the compound of formula (IIa) wherein R^{22} is
cyclopropylmethyl and R^{23} is 4-picolinyl;

the compound of formula (IIa) wherein R^{22} is
benzyl and R^{23} is cyclopentylmethyl;

25 the compound of formula (IIa) wherein R^{22} is
cyclopropylmethyl and R^{23} is cyclopentylmethyl;

the compound of formula (IIa) wherein R^{22} is
benzyl and R^{23} is n-propyl;

30 the compound of formula (IIa) wherein R^{22} is
cyclopropylmethyl and R^{23} is cinnamyl;

35 the compound of formula (IIa) wherein R^{22} is
cyclopropylmethyl and R^{23} is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is cyclopentylmethyl and R²³ is 2-naphthylmethyl;

5 the compound of formula (IIa) wherein R²² is benzyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 2-picolinyl;

10 the compound of formula (IIa) wherein R²² is 3-cyanobenzyl and R²³ is 3-cyanobenzyl;

the compound of formula (IIa) wherein R²² is 3-allyl and R²³ is 2-naphthylmethyl;

15 the compound of formula (IIa) wherein R²² is n-propyl and R²³ is 2-naphthylmethyl;

20 the compound of formula (IIa) wherein R²² is n-butyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is H and R²³ is 2-naphthylmethyl;

25 the compound of formula (IIa) wherein R²² is 4-picolinyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is 3-allyl and R²³ is cyclopentylmethyl;

30 the compound of formula (IIa) wherein R²² is 3-allyl and R²³ is 2-quinolinylmethyl;

35 the compound of formula (IIa) wherein R²² is 3-picolinyl and R²³ is cyclopropylmethyl;

the compound of formula (IIa) wherein R²² is 3-picolinyl and R²³ is 2-naphthylmethyl;

5 the compound of formula (IIa) wherein R²² is 3-allyloxybenzyl and R²³ is 3-allyloxybenzyl;

the compound of formula (IIa) wherein R²² is 3-allyloxybenzyl and R²³ is 3-hydroxybenzyl;

10 the compound of formula (IIa) wherein R²² is 3-picolinyl and R²³ is 3-picolinyl;

the compound of formula (IIa) wherein R²² is 2-naphthylmethyl and R²³ is 4-fluorobenzyl;

15 the compound of formula (IIa) wherein R²² is 3-carbomethoxybenzyl and R²³ is 3-carbomethoxybenzyl;

the compound of formula (IIa) wherein R²² is 4-formylbenzyl and R²³ is 4-formylbenzyl;

20 the compound of formula (IIa) wherein R²² is 4-cyanobenzyl and R²³ is 4-cyanobenzyl;

25 the compound of formula (IIa) wherein R²² is 4-formylbenzyl and R²³ is 4-formylbenzyl;

the compound of formula (IIa) wherein R²² is 4-cyanobenzyl and R²³ is 4-cyanobenzyl;

30 the compound of formula (IIa) wherein R²² is 4-formylbenzyl and R²³ is 4-formylbenzyl;

the compound of formula (IIa) wherein R²² is 4-hydroxybenzyl and R²³ is n-propyl;

35

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the compound of formula (IIa) wherein R²² is 3-hydroxybenzyl and R²³ is n-propyl;

5 the compound of formula (IIa) wherein R²² is 3-carboxybenzyl and R²³ is 3-carboxybenzyl;

the compound of formula (IIa) wherein R²² is cyclobutylmethyl and R²³ is cyclobutylmethyl;

10 the compound of formula (IIa) wherein R²² is cyclopentylmethyl and R²³ is cyclopentylmethyl;

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is 3-methylallyl;

15 the compound of formula (IIa) wherein R²² is n-butyl and R²³ is cyclopentylmethyl;

20 the compound of formula (IIa) wherein R²² is 3-formaldoximebenzyl and R²³ is 3-formaldoximebenzyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 3-hydroxybenzyl;

25 the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 4-hydroxybenzyl;

30 the compound of formula (IIa) wherein R²² is 3-(N-methylamino)benzyl and R²³ is 3-(N-methylamino)benzyl;

the compound of formula (IIa) wherein R²² is 3-acetylbenzyl and R²³ is 3-acetylbenzyl;

the compound of formula (IIa) wherein R^{22} is 3-hydroxylaminobenzyl and R^{23} is 3-hydroxylaminobenzyl;

5 the compound of formula (IIa) wherein R^{22} is 2-naphthylmethyl and R^{23} is 3-hydroxybenzyl;

the compound of formula (IIa) wherein R^{22} is 4-hydroxymethylbenzyl and R^{23} is 3-hydroxybenzyl;

10

the compound of formula (IIa) wherein R^{22} is N-methyl-(3-amido)benzyl and R^{23} is N-methyl-(3-amido)benzyl;

15 the compound of formula (IIa) wherein R^{22} is N-methyl-(3-amido)benzyl and R^{23} is 3-(amidino)benzyl;

the compound of formula (IIa) wherein R^{22} is 3-(5-tetrazolyl)benzyl and R^{23} is cyclopropylmethyl;

20

the compound of formula (IIa) wherein R^{22} is 3-(5-tetrazolyl)benzyl and R^{23} is 3-(5-tetrazolyl)benzyl;

25 the compound of formula (IIa) wherein R^{22} is phenylmethyl-3-boronic acid and R^{23} is phenylmethyl-3-boronic acid.

30

The following preferred compounds of formula (I) exhibit IC_{90} values of less than about 10 mg/mL for the inhibition of HIV growth:

35 the compound of formula (IIa) wherein R^{22} is allyl and R^{23} is allyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cyclopropylmethyl;

5 the compound of formula (IIa) wherein R²² is n-butyl and R²³ is n-butyl;

the compound of formula (IIa) wherein R²² is CH₂CH=CH(CH₃)₂ and R²³ is CH₂CH=CH(CH₃)₂;

10 the compound of formula (IIa) wherein R²² is propyl and R²³ is propyl;

the compound of formula (IIa) wherein R²² is i-pentyl and R²³ is i-pentyl;

15 the compound of formula (IIa) wherein R²² is benzyl and R²³ is benzyl;

20 the compound of formula (IIa) wherein R²² is 3-hydroxybenzyl and R²³ is 3-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is 4-hydroxybenzyl and R²³ is 4-hydroxybenzyl;

25 the compound of formula (IIa) wherein R²² is 3-hydroxymethylbenzyl and R²³ is 3-hydroxymethylbenzyl;

30 the compound of formula (IIa) wherein R²² is 4-hydroxymethylbenzyl and R²³ is 4-hydroxymethylbenzyl;

the compound of formula (IIa) wherein R²² is 3-aminobenzyl and R²³ is 3-aminobenzyl;

35 the compound of formula (IIa) wherein R²² is 3-carboxybenzyl and R²³ is 3-carboxybenzyl;

the compound of formula (IIa) wherein R^{22} is 3-formylbenzyl and R^{23} is 3-formylbenzyl

5 the compound of formula (IIa) wherein R^{22} is 3-formaldoximebenzyl and R^{23} is 3-formaldoximebenzyl;

the compound of formula (IIa) wherein R^{22} is 3-(N-methylamino)benzyl and R^{23} is 3-(N-methylamino)benzyl;

10

the compound of formula (IIa) wherein R^{22} is 3-acetylbenzyl and R^{23} is 3-acetylbenzyl;

15 the compound of formula (IIa) wherein R^{22} is 3-hydroxylaminobenzyl and R^{23} is 3-hydroxylaminobenzyl;

the compound of formula (IIa) wherein R^{22} is 2-naphthylmethyl and R^{23} is 3-hydroxybenzyl;

20

the compound of formula (IIa) wherein R^{22} is 4-hydroxymethylbenzyl and R^{23} is 3-hydroxybenzyl;

25 the compound of formula (IIa) wherein R^{22} is N-methyl-(3-amido)benzyl and R^{23} is N-methyl-(3-amido)benzyl;

the compound of formula (IIa) wherein R^{22} is N-methyl-(3-amido)benzyl and R^{23} is 3-(amidino)benzyl;

30

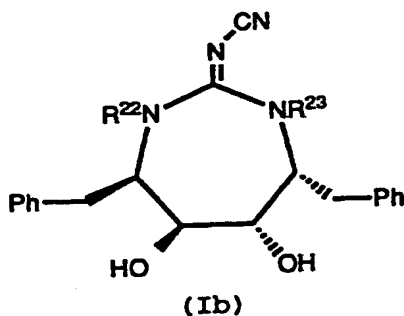
the compound of formula (IIa) wherein R^{22} is 3-(5-tetrazoyl)benzyl and R^{23} is cyclopropylmethyl;

35 the compound of formula (IIa) wherein R^{22} is 3-(5-tetrazolyl)benzyl and R^{23} is 3-(5-tetrazolyl)benzyl;

the compound of formula (IIa) wherein R^{22} is phenylmethyl-3-boronic acid and R^{23} is phenylmethyl-3-boronic acid.

5

Also preferred in the present invention are compounds of formula (Ib):



10

or a pharmaceutically acceptable salt or prodrug form thereof wherein:

15

R^{22} and R^{23} are independently selected from the group consisting of: hydrogen, cyclopropylmethyl, $\text{CH}_2(\text{C}_6\text{H}_4)\text{-p-OCH}_2\text{C}_6\text{H}_5$, $\text{CH}_2(\text{C}_6\text{H}_4)\text{-p-OH}$, cyclopentylmethyl, allyl, n-butyl, beta-naphthylmethyl, benzyl, $\text{CH}_2(\text{C}_6\text{H}_4)\text{-m-OCH}_2\text{C}_6\text{H}_5$, p-nitrobenzyl, m-nitrobenzyl, $\text{CH}_2(\text{C}_6\text{H}_4)\text{-m-OH}$, p-aminobenzyl, m-aminobenzyl, p-nitrilobenzyl, m-nitrilobenzyl, dimethylallyl, cyclohexylmethyl, cyclobutylmethyl, propyl, 3-methyl-1-butyl, carboxamidobenzyl, and formaldoximebenzyl.

20

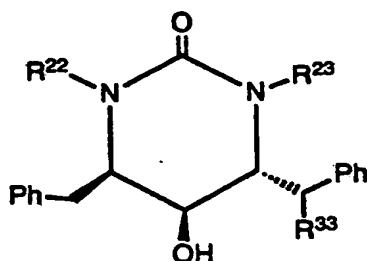
25

More specifically preferred are the following compounds of formula (I), which have IC_{50} or K_i values of less than about 10 nM for inhibiting HIV protease:

30

the compound of formula (Ib) wherein R^{22} is 3-formaldoximebenzyl and R^{23} is 3-formaldoximebenzyl.

- 5 Also preferred in the present invention are compounds of the formula:



- 10 or a pharmaceutically acceptable salt or prodrug form thereof wherein:

R^{33} is OH, halogen, H, N_3 ;

- 15 R^{22} and R^{23} are independently selected from the group consisting of:

hydrogen, allyl, propyl, cyclopropylmethyl,
 n-butyl, i-butyl, $CH_2CH=CH(CH_3)_2$, pyridylmethyl,
 20 methallyl, n-pentyl, i-pentyl, hexyl, benzyl,
 pyridylmethyl, isoprenyl, propargyl, picolinyl,
 methoxyethyl, cyclohexylmethyl, dimethyl-butyl,
 ethoxyethyl, methyl-oxazolinylmethyl,
 naphthylmethyl, methyloxazolinylmethyl,
 25 vinyloxyethyl, pentafluorobenzyl, quinolinylmethyl,
 carboxybenzyl, chloro-thienyl, picolinyl,
 benzyloxybenzyl, phenylbenzyl, adamantylethyl,
 cyclopropylmethoxybenzyl, ethoxybenzyl,
 hydroxybenzyl, hydroxymethylbenzyl, aminobenzyl,
 30 formylbenzyl, cyanobenzyl, cinnamyl,

allyloxybenzyl, fluorobenzyl, cyclobutylmethyl, formaldoximebenzyl, cyclopentylmethyl, nitrobenzyl, nitrilobenzyl, carboxamidobenzyl, carbomethoxybenzyl, and dimethylallyl.

5

In the present invention it has been discovered that the compounds above are useful as inhibitors of HIV protease and similar retroviral proteases, and for the treatment of HIV infection and similar retrovirus infections.

10 The present invention also provides methods for the treatment of HIV infection by administering to a host infected with HIV a pharmaceutically effective amount of a compound of formula (I) as described above.

15

The compounds herein described may have asymmetric centers. All chiral, diastereomeric, and racemic forms are included in the present invention. Many geometric isomers of olefins, C=N double bonds, and the like can also be present in the compounds described herein, and all such stable isomers are contemplated in the present invention.

20 When any variable (for example, R¹ through R²⁸, R^{4A} and R^{7A}, m, n, W, Z, etc.) occurs more than one time in any constituent or in formula (I) or (II), or any other formula herein, its definition on each occurrence is independent of its definition at every other occurrence. Also, combinations of substituents and/or variables are permissible only if such combinations result in stable compounds.

30 As used herein, "alkyl" is intended to include both branched and straight-chain saturated aliphatic hydrocarbon groups having the specified number of carbon atoms; "alkoxy" represents an alkyl group of indicated number of carbon atoms attached through an oxygen bridge; "cycloalkyl" is intended to include saturated

ring groups, such as cyclopropyl, cyclobutyl, cyclopentyl, cyclohexyl, cycloheptyl and cyclooctyl; and "biycloalkyl" is intended to include saturated bicyclic ring groups such as [3.3.0]bicyclooctane, [4.3.0]bicyclononane, [4.4.0]bicyclodecane (decalin), [2.2.2]bicyclooctane, and so forth. "Alkenyl" is intended to include hydrocarbon chains of either a straight or branched configuration and one or more unsaturated carbon-carbon bonds which may occur in any stable point along the chain, such as ethenyl, propenyl, and the like; and "alkynyl" is intended to include hydrocarbon chains of either a straight or branched configuration and one or more triple carbon-carbon bonds which may occur in any stable point along the chain, such as ethynyl, propynyl and the like. "Halo" as used herein refers to fluoro, chloro, bromo, and iodo; and "counterion" is used to represent a small, negatively charged species such as chloride, bromide, hydroxide, acetate, sulfate, and the like.

As used herein, "aryl" or "aromatic residue" is intended to mean phenyl or naphthyl; "carbocyclic" is intended to mean any stable 5- to 7- membered monocyclic or bicyclic or 7- to 14-membered bicyclic or tricyclic carbon ring, any of which may be saturated, partially unsaturated, or aromatic, for example, indanyl or tetrahydronaphthyl (tetralin).

As used herein, the term "heterocycle" is intended to mean a stable 5- to 7- membered monocyclic or bicyclic or 7- to 10-membered bicyclic heterocyclic ring which is either saturated or unsaturated, and which consists of carbon atoms and from 1 to 3 heteroatoms selected from the group consisting of N, O and S and wherein the nitrogen and sulfur heteroatoms may optionally be oxidized, and the nitrogen may optionally be quaternized, and including any bicyclic group in which any of the above-defined heterocyclic rings is

fused to a benzene ring. The heterocyclic ring may be attached to its pendant group at any heteroatom or carbon atom which results in a stable structure. The heterocyclic rings described herein may be substituted on carbon or on a nitrogen atom if the resulting compound is stable. Examples of such heterocycles include, but are not limited to, pyridyl, pyrimidinyl, furanyl, thienyl, pyrrolyl, pyrazolyl, imidazolyl, tetrazolyl, benzofuranyl, benzothiophenyl, indolyl, indolenyl, quinolinyl, isoquinolinyl or benzimidazolyl, piperidinyl, 4-piperidonyl, pyrrolidinyl, 2-pyrrolidonyl, pyrrolinyl, tetrahydrofuranyl, tetrahydroquinolinyl, tetrahydroisoquinolinyl, decahydroquinolinyl or octahydroisoquinolinyl. The term "substituted", as used herein, means that an one or more hydrogen on the designated atom is replaced with a selection from the indicated group, provided that the designated atom's normal valency is not exceeded, and that the substitution results in a stable compound.

By "stable compound" or "stable structure" is meant herein a compound that is sufficiently robust to survive isolation to a useful degree of purity from a reaction mixture, and formulation into an efficacious therapeutic agent.

As used herein, "pharmaceutically acceptable salts and prodrugs" refer to derivatives of the disclosed compounds that are modified by making acid or base salts, or by modifying functional groups present in the compounds in such a way that the modifications are cleaved, either in routine manipulation or *in vivo*, to the parent compounds. Examples include, but are not limited to, mineral or organic acid salts of basic residues such as amines; alkali or organic salts of acidic residues such as carboxylic acids; acetate, formate and benzoate derivatives of alcohols and amines; and the like.

Pharmaceutically acceptable salts of the compounds of the invention can be prepared by reacting the free acid or base forms of these compounds with a stoichiometric amount of the appropriate base or acid in water or in an organic solvent, or in a mixture of the two; generally, nonaqueous media like ether, ethyl acetate, ethanol, isopropanol, or acetonitrile are preferred. Lists of suitable salts are found in Remington's Pharmaceutical Sciences, 17th ed., Mack Publishing Company, Easton, PA, 1985, p. 1418, the disclosure of which is hereby incorporated by reference.

Synthesis

The compounds of the present invention may be synthesized using the general synthetic procedures described below. Each of the references cited below are hereby incorporated herein by reference.

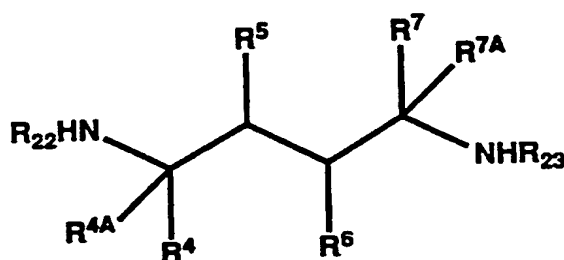
Compounds of the invention wherein:

W is $-N(R^{22})C(=Z)N(R^{23})-$ or
 $-N(R^{22})C(=O)C(=O)N(R^{23})-$;

R^5 is $-OR^{20}$ or H;

R^6 is $-OR^{21}$ or H; and

n is 1; can be formed from diamines of formula (III):



(III)

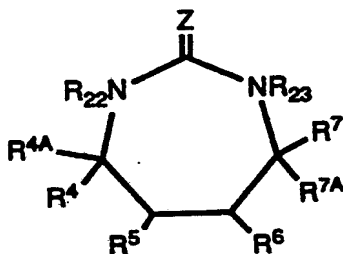
5 The diamines of formula (III) can be synthesized as described in copending commonly assigned patent application Jadhav et al. USSN 07/714,042, filed 5/31/91. Alternative methods which can be used to synthesize the compounds of structure (III) above are
 10 described in European Patent Application Publication Number 402646A1, U.S. Patent 4,837,204, and Canadian Patent Application 2,026,832 .

The compounds of formula (III) can be cyclized to
 15 form compounds of formula (IV) under conditions normally used to form cyclic ureas, as is known to one skilled in the art. Reagents J-(C=Z)-J', where J and J' are leaving groups, are employed, preferably under relatively dilute conditions (for example, less than
 20 about 0.1 M), to effect ring closure to provide compounds of formula (I). Many examples of J and J' are known; preferred are carbonyl diimidazole, thiocarbonyldiimidazole, phosgene, thiophosgene, diphenyl carbonate, or diphenyl thiocarbonate.
 25 Additionally, for compounds wherein W is -N(R²²)C(=O)C(=O)N(R²³)-, compounds of formula (III) can be reacted with activated derivatives of oxalic acid, preferably oxalyl chloride, under the above conditions to form the diamide.

30

For compounds of the invention where R²⁰ or R²¹ is -OH, it is advantageous to protect the free hydroxyl before cyclization. Protecting groups used can include any of those listed in Greene, Protective Groups in Organic Synthesis, Chapter 2, Wiley, NY (1981). The preferred protecting groups are trimethylsilylethoxymethyl (SEM), methoxyethoxymethyl (MEM), or methoxymethyl (MOM).

- 10 Cyclization of compounds of formula (III) results in structure (IV) (i.e., structure (I) wherein W is -N(R²²)C(=O)C(=O)N(R²³)- and n is 1).



15

(IV)

- Another, preferred method to form compounds of formula (IV), in cases wherein R²² and R²³ are linked to their respective nitrogens by a CH₂ residue, is to cyclize a compound of structure (III) where R²² and R²³ are hydrogen, and to alkylate the nitrogens using a base, a phase transfer catalyst, and an alkylating agent, using methods well known in the art. The preferred base is sodium hydride, and the preferred alkylating agents are R²²Y and R²³Y, wherein Y is a halogen, triflate, or mesylate, preferably a bromide or iodide. Preferred conditions are in polar aprotic solvents between 0 and 100 °C.

30

Cleavage of protecting groups, if employed, yields structures of formula (I) wherein R⁵ and R⁶ are hydroxyl.

5 When R⁵ and R⁶ are other than OR²⁰ and OR²¹, some chemical manipulation of functional groups may need to be performed in the preparation of the compounds of formula (III) or (IV), as is appreciated by one of skill in the art of organic synthesis. Described below are
10 examples of such procedures.

15 Methods for obtaining compounds wherein R⁵ is OH and R⁶ is OR²¹ include protection of nitrogen, if necessary, followed by reaction of the diol with one equivalent of base and one equivalent of acyl halide, alkyl halide, alkoxyalkyl halide, alkoxy carbonyl halide, benzoyl halide, diphenyl carbonate or phenylisocyanate, and purification by column chromatography of the unwanted bis-alkylated
20 and unreacted material.

25 Methods for obtaining compounds wherein R⁵ is OH and R⁶ is H include protection of nitrogen, if necessary, and reduction of the diol to the monool using techniques known in the art (see, for example, *Chem. Comm.* 1971, 1097; *J. Org. Chem.* 1969, 3923). The preferred method is formation of cyclic diol ester and reduction using hydride. Deprotection of nitrogen, if necessary, results in
30 the desired compound.

35 Methods for obtaining compounds wherein R⁵ is OH and R⁶ is F include protection of nitrogen, if necessary, followed by formation of mono-protected diol as described above. Reaction with a fluorinating agent, preferably

diethylaminosulfurtrifluoride (DAST) (Reagents for Organic Synthesis, Vol. 13, p. 110, Wiley Interscience, NY, 1988), provides the alkyl fluoride. Deprotection of nitrogen, if necessary, and hydroxyl results in the desired compound.

Methods for obtaining compounds wherein R^5 is OH and R^6 is =O include protection of nitrogen, if necessary, and standard conditions for oxidizing glycols to pinacols. The preferred oxidant is one equivalent of pyridinium dichromate in dichloromethane, or one equivalent of NaOCl in HOAc. Deprotection of nitrogen, if necessary, results in the desired compound. Alternatively, a monohydroxy compound described above can be oxidized to the ketone under standard conditions, preferably Swern oxidation using oxalyl chloride, DMSO and Et₃N, followed by alpha-hydroxylation of the ketone (see Tet. Lett. 1981, 607; Tet. Lett. 1982, 2917).

Methods for obtaining compounds wherein R^5 is OH and R^6 is difluoro include protection of nitrogen, if necessary, and hydroxyl of the above obtained pinacol, followed by reaction of the carbonyl with a fluorinating reagent, such as DAST. Deprotection of hydroxyl and nitrogen, if necessary, results in the desired compound.

Methods for obtaining compounds wherein R^5 and R^6 join to form an epoxide include protection of nitrogen, if necessary, followed by standard conditions for the formation of an epoxide from a glycol (see, for example, J. Org. Chem. 1981, 3361). Preferred is the reaction of the glycol

with more than 2 equivalents of base and one equivalent of an activating group, such as methanesulfonyl chloride. Deprotection if necessary results in the desired compound.

5 Methods for obtaining compounds wherein R^5 is OH and R^6 is C₁-C₃ alkyl include protection of nitrogen, if necessary, and reaction of the epoxide prepared above with C₁-C₃ alkylmetal reagents.
 10 Preferred is the reaction of lithium dialkyl cuprates in aprotic solvents at low temperatures (-78 to -40 °C) (see Carruthers, Some Modern Methods in Organic Synthesis, p. 64, Cambridge University Press, 1978).

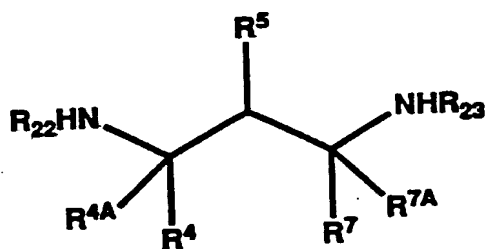
15 With a judicious selection of reagents, as is well appreciated to one skilled in the art, these manipulations can be performed in a straightforward manner to yield the claimed combinations of R^5 and R^6 .

20

Compounds of the invention wherein:

25 W is $-N(R^{22})C(=Z)N(R^{23})-$ or $-N(R^{22})C(=O)C(=O)N(R^{23})-$, and n is 0;

can be synthesized from diamines of formula (V):



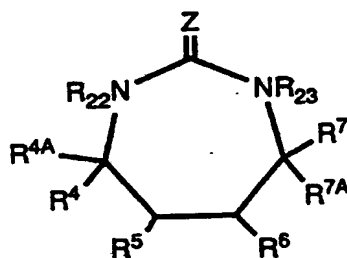
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(V)

which can in turn be synthesized as described in
European Patent Application Publication Number 402 646
A1.

5

Protection, if necessary, cyclization, and
functional group manipulation if desired is performed as
described above to obtain compounds of structure (VI):



10

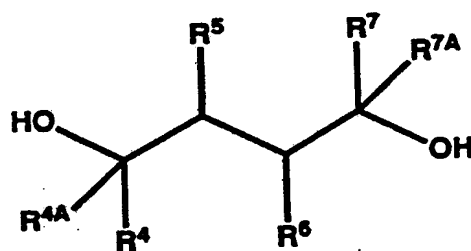
(VI)

Compounds of the invention wherein:

15

W is $-\text{OC}(=\text{O})\text{O}-$ and n is 1;

can be formed from diols of structure (VII):



20

(VII)

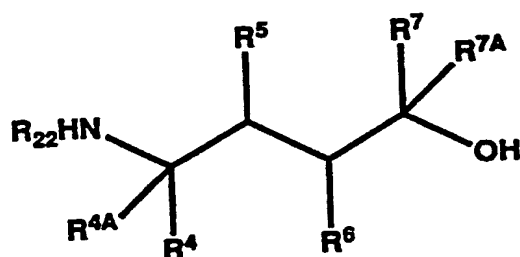
which can in turn be synthesized as described in
25 copending, commonly assigned U.S. patent application
Jadhav et al. USSN 07/714,042, filed 5/31/91.

Functional group manipulation, if desired, may be performed as described above, followed by cyclization to the carbonate using standard conditions, preferably phosgene or thiophosgene in the presence of 2 equivalents of a base such as potassium hydride, to obtain compounds of structure (I).

Compounds of the invention wherein:

W is $-N(R^{22})C(=Z)O-$ and n is 1;

can be formed from aminoalcohol of structure (VIII):



(VIII)

which can in turn be synthesized as described in a copending, commonly assigned U.S. patent application Jadhav et al. USSN 07/714,042, filed 5/31/91, by employing a single equivalent of azide in the reaction of the diol of formula (VII) to obtain the azidoalcohol, followed by reduction as described in USSN 07/714,042, to form the aminoalcohol.

Protection, if necessary, and functional group manipulation, if desired, is performed as described above, followed by cyclization to the carbamate using

standard conditions, preferably phosgene or thiophosgene in the presence of 2 equivalents of a base, such as potassium hydride, to obtain compounds of structure (I).

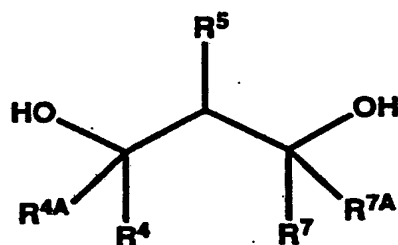
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Compounds of the invention wherein:

W is $-\text{OC}(=\text{Z})\text{O}-$ and n is 0;

10

can be formed from the diol of structure (IX):



(IX)

15

which can in turn be synthesized by the reaction of R^4CHO with the lithium anion of 1,3 dithiane, followed by the reaction of R^7CHO with the anion of the product (see Carruthers, Some Modern Methods in Organic

20

Synthesis, p. 45, Cambridge University Press, 1978). Cleavage of the dithiane with mercuric ion yields the acyclic alpha, alpha' dihydroxyketone.

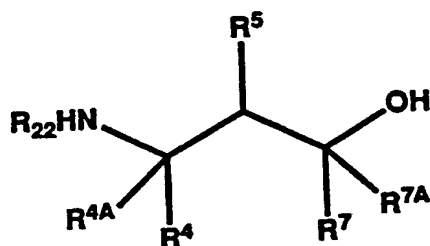
Functional group manipulation, if desired, is performed as described above, followed by cyclization to the carbonate using standard conditions, preferably phosgene or thiophosgene, in the presence of 2 equivalents of a base such as potassium hydride, to obtain compounds of structure (I).

30

Compounds of the invention wherein:

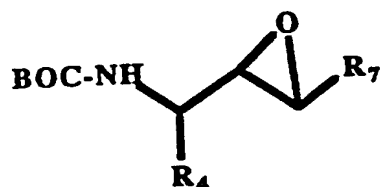
W is $-N(R^{22})C(=Z)O-$ and n is 0;

5 can be formed from aminoalcohol of structure (X):



(X)

10 which can in turn be synthesized by the techniques described in European Patent Application Publication Number 402 646 A1 for the synthesis of compounds of structure (V), above; however, in place of azide, in
 15 opening the oxirane (shown below), an oxygen nucleophile, such as acetate or hydroxide ion, is reacted in the presence of a polar aprotic solvent, such as DMSO.



20

Alternatively, the oxirane is treated with a catalytic amount of a strong acid in water and a cosolvent, if necessary, which technique also removes
 25 the BOC protecting group.

Protection, if necessary, and functional group manipulation, if desired, is performed as described above, followed by cyclization to the carbamate using standard conditions, preferably phosgene or thiophosgene in the presence of 2 equivalents of a base such as potassium hydride, to obtain compounds of structure (I).

Compounds of the present invention wherein:

10

W is $-C(R^{25})(R^{26})N(CH_3)(O)C(R^{27})(R^{28})-$;

can be synthesized from aminoalcohols (VIII) and (X) by the following steps: protection of nitrogen, if necessary, preferably with a benzyloxycarbonyl group; activation of the alcohol to displacement, preferably with a sulfonate derivative, such as mesyl chloride; removal of the nitrogen protecting group, preferably with hydrogen in the presence of a catalyst, such as palladium on carbon; and heating under dilute conditions in the presence of a base such as triethylamine to effect cyclization.

The secondary cyclic amine is then methylated, preferably with formic acid/formaldehyde, and oxidized, preferably with a peracid, such as MCPBA, to form compounds of formula (I), wherein W is $-C(R^{25})(R^{26})N(CH_3)(O)C(R^{27})(R^{28})-$. The secondary cyclic amine can alternatively be directly oxidized to form structure (I), where W is $-C(R^{25})(R^{26})N(OR^{29})C(R^{27})(R^{28})-$.

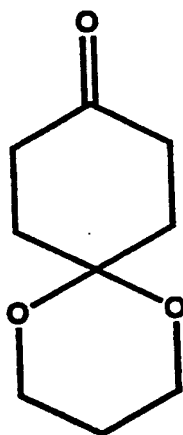
30

Compounds wherein:

35

W is $-C(R^{25})(R^{26})C(=Z)C(R^{27})(R^{28})-$ and n is 0;

can be prepared by the alkylation of protected
 cyclohexanedione (XI) with the required R^4 -LG and R^7 -LG,
 and optionally R^{4A} -LG and R^{7A} -LG groups, wherein LG
 5 represents a leaving group such as halogen or sulfonate
 ester.



(XI)

10

Reduction of the ketone to the alcohol, preferably
 with LiAlH_4 , or manipulation to other values of R^5 as
 described above, is followed by cleavage of the ketal
 15 (see Greene, Protective Groups in Organic Synthesis,
 Chapter 2, Wiley, NY, 1981). Protection of the alcohol
 or other reactive groups, followed by alkylation ketone
 and deprotection, provides compounds of structure (I),
 wherein W is

20 $-\text{C}(\text{R}^{25})(\text{R}^{26})\text{C}(=\text{Z})\text{C}(\text{R}^{27})(\text{R}^{28})-$ and n is 0.

Compounds wherein:

25 W is $-\text{C}(\text{R}^{25})(\text{R}^{26})\text{C}(=\text{Z})\text{C}(\text{R}^{27})(\text{R}^{28})-$ and n is 1;

- can be prepared from the protected hydroxyketones described immediately above by ring expansion, for example via the Tiffeneau-Demyanov reaction (March, Advanced Organic Chemistry, p. 965, Wiley, NY, 1985), or
- 5 by treatment with dimethylsulfonium ylide to form the spiro-epoxide, followed by acid-catalyzed ring expansion to the cycloheptanone (ibid., pp. 871, 966).

- The above routes have the advantage of producing a
- 10 number of stereoisomers which, upon purification, can be evaluated for the best combination of potency, safety and in vivo availability.

- 15 Compounds wherein:

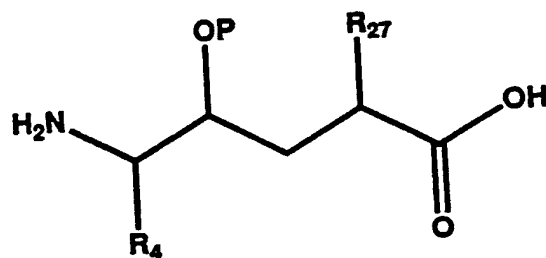
W is $-C(R^{25})(R^{26})C(F_2)C(R^{27})(R^{28})-$ and
n is 0 or 1;

- 20 can be obtained from the above-described protected hydroxyketone by treatment with a fluorinating reagent, preferably DAST, as described above.

- 25 Compounds wherein:

W is $-N(R^{22})C(=Z)C(R^{27})(R^{28})-$ and n is 0;

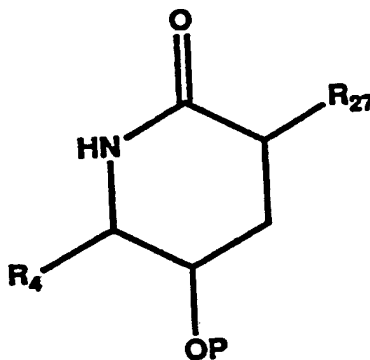
- 30 can be obtained by cyclization of compound (XII) to the lactam using techniques known in the art (March, Advanced Organic Chemistry, p. 371, Wiley, NY, 1985).



(XII)

5 Compounds of structure (XII) can in turn be
obtained as described in European Patent Application
Publication Number 434 365 A2, European Patent
Application Publication Number 386 611 A2, European
Patent Application Publication Number 389 127 A1, and CA
10 2005337, each of which are hereby incorporated by
reference. OP in structure (XII) designates protected
oxygen. Hydroxyl can be protected by the use of any of
a number of groups as described in Greene, Protective
Groups in Organic Synthesis, Chapter 2, Wiley, NY
15 (1981).

If desired, the resulting lactam (XIII):



(XIII)

can be further functionalized, for example by the following techniques: the lactam nitrogen can be alkylated with an R^{22} -LG group, preferably employing sodium hydride in DMF; an R^{4A} , R^7 or R^{7A} group can be added by deprotection and oxidation of the alcohol, followed by alkylation of the enolate using R^{4A} -LG, R^7 -LG or R^{7A} -LG; and reduction of the ketone to hydroxyl or otherwise functionalizing to obtain the R^5 group of choice as described above.

10

Compounds wherein:

W is $-N(R^{22})C(=Z)C(R^{27})(R^{28})-$ and n is 1;

15

can be obtained through techniques known in the art from ketones of structure (I) wherein W is $-C(R^{25})(R^{26})C(=Z)C(R^{27})(R^{28})-$ and n is 0, preferably via the Beckmann rearrangement (March, Advanced Organic Chemistry, p. 987, Wiley, NY, 1985). Manipulation of the R^5 group, if desired, as described above provides R^5 and R^6 -substituted examples of (I), wherein W = $-N(R^{22})C(=Z)C(R^{27})(R^{28})-$ and n = 1.

20

25 Compounds wherein:

W is $-C(R^{25})(R^{26})C(=Z)O-$ and n is 0 or 1;

can be obtained from compounds of structure (I), wherein W = $-N(R^{22})C(=Z)C(R^{27})(R^{28})-$, n = 0 or 1, and $R^{22} = H$, for example, by hydrolysis of the lactam, followed by displacement of the primary amine by hydroxyl, and closure to the lactone (March, Advanced Organic Chemistry, p. 348, Wiley, NY, 1985).

30

Similarly, compounds wherein:

W is $-C(R^{25})(R^{26})C(=Z)S-$ and n is 0 or 1;

5 can be obtained from compounds of structure (I), wherein
W = $-N(R^{22})C(=Z)C(R^{27})(R^{28})-$, n = 0 or 1, and $R^{22} = H$,
for example, by hydrolysis of the lactam, followed by
conversion of the primary amine to the diazonium salt,
displacement by NaSH, and closure to the thiolactone
(March, Advanced Organic Chemistry, p. 601, Wiley, NY,
10 1985).

Compounds of structure (I) described above wherein
Z = O can be converted to the thio derivatives, Z = S,
using standard conditions (March, Advanced Organic
15 Chemistry, p. 792, Wiley, NY, 1985); preferred is the
use of the disulfide described in Bull. Soc. Chim.
Belges 1978, 223.

Structures (I) described above wherein Z = O can be
20 converted to the imino derivatives, Z = NR^{24} , using
standard conditions. When R^{24} is OH or O-alkyl, the
oximes can be formed and alkylated if desired as
described in March, Advanced Organic Chemistry, pp. 359,
805, Wiley, NY, 1985. The hydrazones and imines can be
25 formed similarly (ibid, pp. 533, 797).

It is expected that the compounds of the invention
can also be prepared as shown in Scheme 1 (shown below).
The intra-molecular coupling of the N-substituted or
30 unsubstituted dialdehydes may be achieved by organometal
reagents derived from vanadium, titanium, samarium etc.
The dialdehyde precursors can be prepared from the
commercially available materials by the methods known to
those skilled in the art of organic synthesis,
35 preferably by the techniques disclosed in copending

commonly assigned U.S. Patent Application, Hodge, USSN 07/659,442, filed 2/21/91.

5 Compounds wherein W is $-N(R^{22})C(=O)N(R^{23})-$ and n is 2 can be synthesized as shown in Scheme 2 (below). The eight-membered cyclic urea in Scheme 2 can be protected, if necessary, and manipulated as described above to yield the desired compounds.

10 Compounds wherein W is $-N(R^{22})C(=O)N(R^{23})-$ and n is 1 can likewise be synthesized as shown in Schemes 3, 4, 6, 7 (below). If necessary, intermediates described herein can be manipulated by methods known to those skilled in the art of organic synthesis to provide
15 compounds within the scope of the invention.

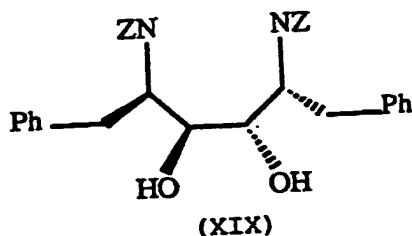
20 Compounds wherein W is $-N(R^{22})C(=N-OR)N(R^{23})-$ or $-N(R^{22})C(=S)N(R^{23})-$ and n is 1 can be synthesized as shown in Scheme 5 (below). If necessary, intermediates described herein can be manipulated by methods known to those skilled in the art of organic synthesis to provide
25 compounds within the scope of the invention.

30 Compounds wherein W is $-N(R^{22})C(=O)N(R^{23})-$ and n is 0 can likewise be synthesized as shown in Scheme 8 (below). If necessary, intermediates described herein can be manipulated by methods known to those skilled in the art of organic synthesis to provide compounds within the scope of the invention.

35 The synthesis of compounds of the invention is described in further detail below.

Procedure 1

Preparation of di-N-CBZ protected 1,4-diamino-2,3-diols (XIX):



5

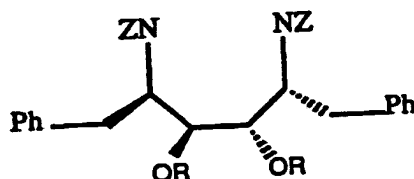
Detailed experimental procedures for the synthesis of compound (XIX) are described in copending commonly assigned patent application Jadhav et al. USSN 07/714,042, filed 5/31/91.

10

Procedure 2

Preparation of di-O protected di-N-CBZ 1,4-diamino diols (XXa) and (XXb):

15



(XXa) R=SEM
(XXb) R=MOM

20

A. Protection as 2-(trimethylsilyl) ethoxy methyl (SEM) ether (XXa):

25

Compound (XIX) (60 g, 105 mmol) was dissolved in dry DMF (600 mL). Diisopropylethylamine (75 mL) and SEMCl (66.8 g, 400mmol) were added and the mixture stirred for 16 h at room temperature under N₂. The solution was diluted with water (1L) and extracted with

hexane (400mL). The organic layer was separated and washed with water (2x100mL). The aqueous layers were combined and extracted with hexane (2x300mL). The organic layers were combined, washed with water (2x100mL), dried over MgSO_4 , filtered and evaporated. The residue was chromatographed on SiO_2 and eluted with 10-30% ethyl acetate/hexane to afford a white solid (91g, 100%). NMR(CDCl_3): δ 7.0-7.4 (m, 20H, Ph), 5.01 (br s, 4H, PhCH_2CO), 4.5-4.95 (m, 6H, NH, OCH_2O), 3.6-4.25 (m, 4H, CHOCH_2 , CHNH), 3.5 (s, 4H, OCH_2CH_2), 2.76 (br d, 4H, PhCH_2), 0.8-1.0 (m, 4H, SiCH_2). MS: 846 ($\text{M}+\text{NH}_4$, 100), 695 (M-SEM, 40).

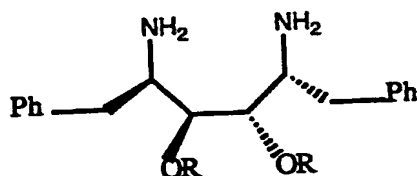
B. Protection as methoxymethyl (MOM) ether (XXb).

Compound (XIX) (.50 g, 0.88 mmol) was dissolved in dry DMF (10mL). Diisopropylethylamine (.46 mL, 2.64 mmol) and methoxymethyl bromide (.165 mL, 2.02 mmol) were added and the solution stirred at 40 °C under nitrogen for four h. TLC (50/50 ethyl acetate /methylene chloride) showed that the reaction was complete. The mixture was partitioned between methylene chloride (50 mL) and 5% HCl (30 mL). The organic layer was separated, washed with water (5x20 mL), brine (20 mL), dried over MgSO_4 , filtered and evaporated to a light yellow oil. Chromatography on SiO_2 and elution with 1-20% ethyl acetate/methylene chloride afforded (XXb) as a clear oil (.29 g, 53%). NMR (CDCl_3): δ 6.95-7.42 (m, 20H, Ph), 5.1-3.8 (m, complex), 3.35 (s, 6H, OCH_3), 2.8-2.95 (m, 4H, PhCH_2). MS: 657 (13, $\text{M}+1$), 674 (21, $\text{M}+\text{NH}_4$), 522 (84), 414 (100), 370 (34).

Procedure 3

35

Deprotection of amines (XXa) and (XXb) via hydrogenation to afford (XXIa) and (XXIb):



5

(XXIa) R=SEM

(XXIb) R=MOM

A. Hydrogenation of SEM ether (XXa).

10

Compound (XXa) (90 g, 108.5 mmol) was dissolved in absolute ethanol (2.5 L). 5% Pd/C (6.5 g) was added and the solution was stirred under hydrogen for 1.5 h until hydrogen uptake ceased. TLC (20/80 ethyl acetate/hexane) showed that the reaction was complete. The solution was filtered through Celite and evaporated at reduced pressure to a colorless gum (60 g, 99%). NMR (CDCl₃): δ 7.1-7.35 (m, 10H, Ph), 4.72 (br d, 4H, OCH₂O), 3.5-3.9 (m, 6H, NH₂, CHOCH₂), 3.15 (m, 2H, CHNH₂), 2.55-2.95 (m, 4H, PhCH₂), 0.95 (m, 4H, SiCH₂).

15

B. Hydrogenation of MOM ether (XXb).

Compound (XXb) (.29 g, .441 mmol) was dissolved in ethyl acetate (6 mL) and methanol (3 mL). 10% Pd/C (70 mg) was added and the solution stirred under hydrogen until H₂ uptake ceased. TLC (20/80 methanol/ethyl acetate) showed that the reaction was complete. The solution was filtered through Celite and evaporated at reduced pressure to afford 3b as a clear oil (.132 g, 77.4%). NMR (CDCl₃): δ 7.1-7.35 (m, 10H, Ph), 4.58 (s, 4H, OCH₂O), 3.75 (br s, 2H, CHOCH₂), 3.3-3.5 (m, 2H,

25

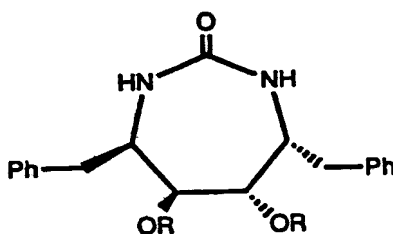
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CHNH₂), 3.23 (s, 6H, OCH₃), 2.85 (br d, 4H, PhCH₂). MS: 389 (M+1, 100), 345 (3.7), 280 (1.8), 120 (6.1).

Procedure 4

5

Formation of cyclic ureas (XXIIa) and (XXIIb):



10

(XXIIa) R=SEM

(XXIIb) R=MOM

A. Cyclization of SEM ether (XXIa).

15 Compound (XXIa) (40 g, 71.3 mmol) was dissolved in methylene chloride (200 mL). Carbonyl diimidazole (13.87 g, 85.6 mmol) was dissolved in methylene chloride (200 mL) in a separate flask. Each solution was then pumped into dry methylene chloride (6 L) at a rate of 90 mL/h.

20 The mixture was then stirred for 18h at room temperature under nitrogen. TLC (60/40 ethyl acetate/hexane) showed the reaction was complete. The solvent was removed at reduced pressure and residue chromatographed on SiO₂ and eluted with 1-50% ethyl acetate/hexane to afford (XXIIa)

25 as a white solid (38.82g, 93%). mp: 75-76°C. NMR (CDCl₃): δ 7.05-7.4 (m, 10H, Ph), 4.6-4.8 (dd, 4H, OCH₂O), 4.08 (s, 2H, CHOCH₂), 3.5-3.91 (m, 8H, NH, CHNH, OCH₂CH₂), 2.86, (br d, 4H, PhCH₂), 0.8-0.95 (m, 4H, SiCH₂). MS: 587 (M+1, 100).

30

B. Cyclization of MOM ether (XXIb).

WO 93/07128

Compound (XXIb) (.53 g, 1.364 mmol) was dissolved in dry methylene chloride (20 mL). In a separate flask, carbonyl diimidazole (.265 g, 1.64 mmol) was dissolved in methylene chloride (20 mL). To a third flask containing pyridine (.22 mL, 2.73 mmol) in methylene chloride (100 mL) at room temperature under nitrogen were added the first two solutions via syringe pump at a rate of 1.7 mL/h. The solution was stirred overnight at room temperature. TLC (50/50 ethyl acetate/methylene chloride) showed that the reaction was complete. The solution was washed with 5% HCl (50 mL), NaHCO₃ (50 mL), brine (50 mL), dried over MgSO₄, filtered and evaporated. The residue was chromatographed on SiO₂ and eluted with 50-75% ethyl acetate/methylene chloride to afford (XXIIb) as a colorless gum (198 mg, 35%). NMR (CDCl₃): δ 7.1-7.4 (m, 10H, Ph), 4.65 (q, 4H, OCH₂O), 4.13 (s, 2H, NH), 3.89 (t, 2H, CHNH), 3.59 (s, 2H, CHOCH₂), 3.18 (s, 6H, OCH₃), 2.87 (m, 4H, PhCH₂). MS: 415 (M+1, 100), 102 (11).

Synthesis of Dimem DiZ Intermediate:

DiZ Diol 507g(0.89mol) was stirred in 4L of dichloromethane. To the slurry was added N,N-Diisopropylethylamine 780g(6.05mol) in one portion at room temperature followed by the dropwise addition of B-methoxyethoxy methylchloride 500g(4mol) (1 hour addition, exothermic). Heated the solution at reflux for 12 hours. TLC (10:1:10 EtOAc:EtOH:Hexanes, R_f=0.56) indicated a complete reaction. The solution was worked up by quenching with ice water(3L). Washed the dichloromethane extract with water(2x 2L) and dried over magnesium sulfate. The filtrate was taken to dryness. The resultant semi-solid was dissolved in chlorobutane(1L). Passed the solution through a four

inch pad of silica gel to remove most of the intense red color. To the chlorobutane extract was added hexane(2L) to precipitate the desired DiZ Dimem intermediate. Washed the white solid with hexanes(3x 350ml). Dried at room temperature. Recovered the desired DiZ Dimem intermediate as a white solid in a yield of 525g(79% yield). m.p. 52-54 C, ¹H NMR(CDCl₃): 2.80(m, 4H)-CH₂Ph, 3.38(s, 6H)-OCH₃, 3.58(m, 8H)-OCH₂CH₂O-, 3.80(m, 2H), 4.20(m, 2H), 4.6-5.2(m, 10H)NH, H₂CCO₂, -OCH₂O-, 7.25(m, 20H)C₆H₅

Synthesis of Cyclic Urea Intermediate:

DiZ Dimem 20g(26.8mmol) was dissolved in 200ml of tetrahydrofuran. To the solution was added 2g of 10% Palladium on Carbon and the suspension stirred for 7 hours under hydrogen(1 atm). TLC (10:1:10 EtOAc:EtOH:Hex, R_f=0.05) indicated a complete reaction. The suspension was filtered through a bed of Celite to remove the catalyst. Washed the Celite bed with 150ml of tetrahydrofuran. Transferred the THF solution to a 500ml round bottom flask. To the THF solution was added 5.5g(33.3mmol) 1,1'-Carbonyldiimidazole in several portions as a solid. Stirred at room temperature for 12 hrs. TLC (10:1:10 EtOAc:EtOH:Hex, R_f=0.26) indicated a complete reaction. The mixture was worked up by quenching with ice-cold 0.5N HCl (150ml) and extracting with diethyl ether (2x50ml). The organic extract was washed with water (2x100ml) and dried over magnesium sulfate. The filtrate was taken to dryness. The residue was purified on silica gel(200g; 1:1 EtOAc:Hex followed by 10:1:10 EtOAc:EtOH:Hex) to provide 10.2g (75.7% yield over two steps) of the desired cyclic urea intermediate as a colorless oil. ¹H NMR(CDCl₃): 2.90(m, 4H)-CH₂Ph, 3.36(s, 6H)-OCH₃, 3.40(m, 8H)-OCH₂CH₂O-, 3.60(m, 2H), 3.90(t, 2H), 4.10(s, 2H)NH, 4.80(q, 4H)-OCH₂O-, 7.30(m, 10H)C₆H₅

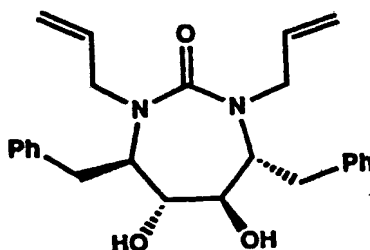
Procedure 5

5 General alkylation/hydrolysis procedure:

Compound (XXIIa) (1 mmol) in dry DMF (5 mL) was added to a flask containing sodium hydride (10 mmol, that had been washed with hexane, 3x20 mL) in DMF (5 mL). The solution was stirred at room temperature under nitrogen for 5 min. Evolution of hydrogen gas was observed. The appropriate alkyl bromide (5 mmol) was added and the solution was stirred at room temperature under nitrogen for 1h. Hindered alkyl bromides required heating at 40-70 °C for up to 5 h. TLC (40/60 ethyl acetate/hexane) was used to ensure that no starting material remained. The solution was quenched with methanol (1 mL), partitioned between ether (60 mL) and water (50 mL) and the organic layer was removed. The aqueous layer was washed with ether (50 mL), the organic layers combined and washed with water (4x30 mL), brine (30 mL), dried over MgSO₄, filtered and evaporated. In cases where the alkyl bromide contained nitrogen, 1 N NaOH was used in place of water.

25 The crude product was hydrolyzed directly in methanol (10 mL) and 4 N HCl/dioxane (5 mL) for up to 16 h at room temperature. The solution was evaporated and chromatographed directly on SiO₂ to afford the bis-alkylated cyclic ureas. Where nitrogen was present, the solutions were first basified with 1 N NaOH and extracted with ethyl acetate, dried over MgSO₄, filtered, evaporated and chromatographed.

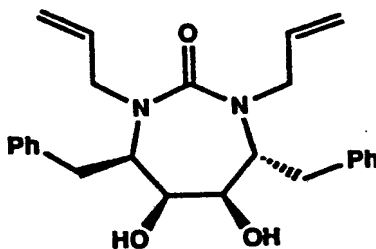
35 Hydrolysis of (XXIIb) under the same conditions gave 67% yield of the N,N-unsubstituted cyclic urea Example 1A, mp 170-174 °C.

Example 1G

5

The experimental procedure is similar to the synthesis of Example 1E. The isomer, (2R,3R,4R,5R)-2,5-diamino-1,6-diphenyl-3,4-hexanediol, needed for the synthesis was isolated from the vanadium trichloride coupling reaction, as described in copending commonly assigned patent application Jadhav et al. USSN 10 07/714,042, filed 5/31/91 (see Procedure 1 above).

Example 1G: ¹³C NMR (CDCl₃) : (75.48 MHz) 37.387, 15 51.51, 65.136, 72.779, 118.649, 126.540, 128.409, 129.714, 134.618, 137.757, 162.705.

Example 1F

20

Compound (XXIIb) (0.85g) was heated with mixture of acetic acid (9.5 mL) and water (0.5 mL) at 85 °C for 4h. After extraction with dichloromethane, followed by washing the organic extract with saturated 25 sodiumbicarbonate and brine, a mixture was provided

which, on separation by column chromatography, furnished (XXIb) (TLC 1:10 ethyl acetate/hexane R_f = 0.4; 0.54g), the desired mono-alcohol intermediate (TLC 1:10 ethyl acetate/hexane R_f = 0.1; 0.13g), and overhydrolysed diol (0.05g).

The above mono-alcohol intermediate 0.25g (0.466 mmol), triphenylphosphine 183mg (0.7 mmol), diethylazadicarboxylate 0.11 mL (0.7 mmol), and chloroacetic acid 66mg (0.7 mmol) were stirred in 5 mL anhydrous tetrahydrofuran at 0 °C for 15 minutes and then at room temperature for 18 h. The excess reagents were quenched with 0.5 mL methanol and the mixture allowed to stir for 20 minutes. The mixture was purified by silica gel column chromatography to provide the desired chloroacetate intermediate with inversion of configuration. ^{13}C NMR (CDCl_3) : (75.48 MHz) -1.373, 14.413, 14.487, 18.253, 25.591, 33.851, 35.741, 40.505, 48.824, 49.962, 57.507, 58.234, 66.589, 67.885, 73.179, 77.423, 95.454, 117.296, 118.554, 126.588, 126.887, 128.518, 128.610, 129.117, 129.199, 129.479, 133.686, 134.168, 136.324, 138.285, 155.698, 166.323.

The above chloroacetate intermediate 73mg (0.12 mmol) in 2mL dry methanol was treated with 0.25 mL (0.5M) sodium methoxide and stirred for 30 minutes at room temperature. The contents were then treated with 0.3 mL (4% HCl in methanol) and stirred for 4.5h at room temperature. The residue after removal of solvent was purified on silica gel column to provide Example 1F.

Example 1F: ^{13}C NMR (CDCl_3): (75.48 MHz) 34.075, 37.672, 48.941, 48.985, 58.071, 60.640, 65.861, 73.212, 177.975, 118.669, 126.535, 126.858, 128.603, 128.815, 129.225, 133.605, 134.172, 137.637, 138.273, 155.497.

35

Synthesis of Monoalkyl Cyclic Urea:

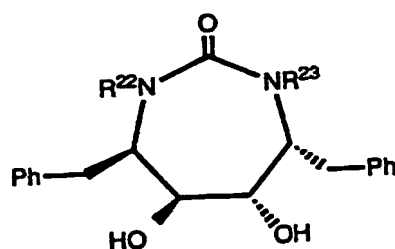
- The intermediate from previous step 2g(4mmol) was dissolved in 25ml toluene and placed in a 100ml round bottom flask. To the solution was added 85% KOH 0.82g(12mmol) and polyethylene glycol (M.W.=1000) 0.20g. With a Dean Stark trap in place the mixture was refluxed for 4 hours until the theoretical amount of water(0.20ml) was collected. Cooled to room temperature and added (bromomethyl)cyclopropane 1.78g(13.2mmol). Stirred at 75 C for 17 hours. TLC(10:1:10 EtOAc:EtOH:Hex, Rf=0.52) indicated that the reaction was complete. Worked up by quenching with aqueous ammonium chloride(50ml) and extracting with ethyl acetate (2x35ml). Washed the organic layer with water(2x35ml) and dried over magnesium sulfate. The filtrate was taken to dryness. The residue was purified on silica gel(150g, 2:3 EtOAc:Hex) to provide 1.55g(70% yield) of the desired monoalkyl cyclic urea as a colorless oil. C13 NMR(CDCl3): 3.331, 4.000, 10.619, 32.877, 34.159, 55.677, 58.294, 58.972, 64.085, 67.361, 67.437, 71.723, 71.753, 76.576, 78.023, 96.347, 96.519, 126.224, 126.316, 128.366, 128.563, 129.400, 129.447, 139.475, 139.555, 161.558.

Examples 1A-1Z, 1AA-1AZ, and 1BA-1BD

25

The compounds listed in Table 1a (Examples 1A-1Z, 1AA-1AZ, and 1BA-1BD) were synthesized using the above-described procedures.

Table 1a

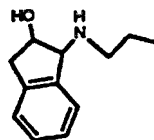
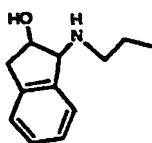


5

Example Number	Stereo-isomer (2,3,4,5)	R ²²	R ²³	Ki
1A	RSSR	H	H	+
1B	SRRS	H	H	+
1C	RSSR	CH ₂ CH=CH ₂	CH ₂ CH=CH ₂	+++
1D	SRRS	CH ₂ CH=CH ₂	CH ₂ CH=CH ₂	+
1E	SSSS	H	H	++
1F	RRSR	CH ₂ CH=CH ₂	CH ₂ CH=CH ₂	++
1G	RRRR	CH ₂ CH=CH ₂	CH ₂ CH=CH ₂	+
1H	SSSS	CH ₂ CH=CH ₂	CH ₂ CH=CH ₂	+
1I	SRRS	CH ₂ Ph	CH ₂ Ph	+
1J	SRRS	(CH ₂) ₂ O(CH ₂) ₂ OMe	(CH ₂) ₂ O(CH ₂) ₂ OMe	+
1K	SRRS	1-geranyl	1-geranyl	+
1L	SRRS	CH ₂ CH ₂ OH	CH ₂ CH ₂ NHCH(CH ₂ CH ₃) ₂	+
			-C(=O)O-benzyl	
1M	SRRS	CH ₂ CH ₂ (CH ₂ CH ₂ O) ₂ OCH ₃	CH ₂ CH ₂ (CH ₂ CH ₂ O) ₂ OCH ₃	+
1N	SRRS	3,7-dimethyl-1-octyl	3,7-dimethyl-1-octyl	++
1O	RSSR	CH ₃ OCH ₂ CH ₂ OCH ₂ CH ₂	CH ₃ OCH ₂ CH ₂ OCH ₂ CH ₂	+
1P	RSSR	3,7-dimethyl-1-octyl	3,7-dimethyl-1-octyl	++
1Q	RSSR	3,7-dimethyl-1-octyl	H	++
1R	RSSR	CH ₃ (OCH ₂ CH ₂) ₃	H	+
1S	RSSR	CH ₃ (OCH ₂ CH ₂) ₃	CH ₃ (OCH ₂ CH ₂) ₃	+
1T	RSSR	HOCH ₂ CH ₂	CH ₂ CH ₂ NHCH(CH ₂ CH ₃) ₂	++
			-C(=O)O-benzyl	

1U	SRRS	CH ₂ CH ₂ NHCH(CH ₃) ₂	CH ₂ CH ₂ NHCH(CH ₃) ₂	+
		-C(=O)O-benzyl	-C(=O)O-benzyl	
1V	RSSR	propyl	propyl	+++
1W	RSSR	methyl	methyl	+
1X	RSSR	cyclopropyl-methyl	cyclopropyl-methyl	+++
1Y	RSSR	n-hexyl	n-hexyl	+++
1Z	RSSR	n-butyl	n-butyl	+++
1AA	RSSR	N-morpholinoethyl	N-morpholinoethyl	+
1AB	RSSR	n-heptyl	n-heptyl	++
1AC	RSSR	CH ₂ CH=C(CH ₃) ₂	CH ₂ CH=C(CH ₃) ₂	+++
1AD	RSSR	ethyl	ethyl	++
1AE	RSSR	CH ₂ CH=C(CH ₃) ₂	H	+++
1AF	RSSR	propyl	propyl	+++
1AG	RSSR	CH ₃	CH ₃	+
1AH	RSSR	i-pentyl	i-pentyl	+++
1AI	RSSR	4-pyridylmethyl	4-pyridylmethyl	++
1AJ	RSSR	2-methallyl	2-methallyl	+++
1AK	RSSR	n-pentyl	n-pentyl	+++
1AL	RSSR	i-octyl	i-octyl	++
1AM	RSSR	i-hexyl	i-hexyl	+++
1AN	RSSR	i-heptyl	i-heptyl	++
1AO	RSSR	i-butyl	i-butyl	++
1AP	RSSR	3-propargyl	3-propargyl	++
1AQ	RSSR	benzyl	benzyl	+++
1AR	RSSR	2-pyridylmethyl	2-pyridylmethyl	++
1AS	RSSR	phenylpropyl	phenylpropyl	++
1AT	RSSR	allyl	isoprenyl	+++
1AU	RSSR	1-cinnamyl	1-cinnamyl	+++
1AV		methyl	cyclopropyl-methyl	
1AW		allyl	cyclopropyl-methyl	
1AX		CH ₂ CH ₂ N(CH ₂) ₂	allyl	
1AY		methyl	benzyl	
1AZ		N-pyrrolylethyl	N-pyrrolylethyl	

1BA



1BB

2-furylmethyl
CH₂CH₂OCH(CH₂)₂

2-furylmethyl
CH₂CH₂OCH(CH₂)₂

1BC

isoprenyl

methyl

1BD

The K_i values in Table 1 were determined using the assay conditions described below under HIV Protease Inhibition Assay. The K_i values are indicated as follows: +++ = <10 nM; ++ = 10 nM to 1 μM; + = >1 μM.

Listed below are physical data for representative compounds of the invention.

10

Example 1W: mp 170–174 °C (67% yield). MS: 355 (M+1, 100). NMR (CDCl₃) δ 7.1–7.35 (m, 10H, Ph), 4.03 (s, 2H, CHOH), 3.5 (d, 2H, NCH), 2.8–3.1 (m, 6H, PhCH₂, OH), 2.58 (s, 6H, NH₃).

15

Example 1AD: mp 214–215 °C. MS: 383 (M+1, 100). NMR (CDCl₃) δ 7.1–7.35 (m, 10H, Ph), 3.95 (s, 2H, CHOH), 3.5 (d, 2H, NCH), 2.8–3.1 (m, 4H, PhCH₂, OH), 2.7 (br s, 2H, OH), 2.3 (m, 2H, NCH), .95 (t, 6H, CH₃).

20

Example 1AF: mp 180–182 °C. MS: 411 (M+1, 100). NMR (CDCl₃) δ 7.05–7.3 (m, 10H, Ph), 4.0 (s, 2H, CHOH), 3.68 (m, 2H, NCH₂), 3.52 (d, 2H, NCH), 3.05 (m, 4H, PhCH₂), 2.1 (m, 2H, NCH₂), 1.6 (s, 2H, OH), 1.4 (m, 4H, MeCH₂),

25

.79 (t, 6H, CH₃).

Example 1Z: HRMS: Calc. 439.2960. Found: 439.2959.

Example 1AK: mp 125-127 °C. MS: 467(M+1, 100).
NMR(CDCl₃) δ 7.15-7.35(m, 10H, Ph), 3.95(s, 2H, CHOH),
3.68(m, 2H, NCH₂), 3.55(d, 2H, NCH), 2.9-3.15(m, 4H,
PhCH₂), 2.75(s, 2H, OH), 2.18(m, 2H, NCH₂), >8-1.45(m,
5 complex, pentyl).

Example 1Y: mp 110-112 °C. MS: 495(M+1, 100). NMR(CDCl₃) δ
7.1-7.35(m, 10H, Ph), 4.0(s, 2H, CHOH), 3.65(m, 2H, NCH-
H), 3.52(d, 2H, NCH), 2.8-3.2(m, 4H, PhCH₂), 2.15(m, 2H,
10 OH), .9-1.45(m, 11H, hexyl).

Example 1AB: mp 100-101 °C. MS: 523(M+1, 100). NMR(CDCl₃) δ
7.1-7.35(m, 10H, Ph), 3.95(s, 2H, CHOH), 3.65(m, 2H,
NCH-H), 3.5(d, 2H, NCH), 2.9-3.1(m, 4H, PhCH₂), 2.6(s,
15 2H, OH), 2.15(m, 2H, NCH), .8-1.4(m, complex, heptyl).

Example 1I: MS: 507(M+1, 100). NMR(CDCl₃) δ 7.05-7.4(m,
20H, Ph), 4.91(d, 4H, PhCH₂N), 3.5-3.65(m, 4H, NCH,
CHOH), 3.05(m, 4H, PhCH₂), 2.35(br s, 2H, OH).

Example 5i (R²² and R²³ = allyl; R²⁰ and R²¹ =
C(=O)CH₃): mp 164-166 °C. MS: 407(M+1, 100). NMR(CDCl₃) δ
7.1-7.4(m, 10H, Ph), 5.65(m, 2H, CH₂CH), 5.01(m, 4H,
CH₂CH), 4.26(m, 4H, NCH₂), 3.91(s, 2H, CHOH), 3.59(d,
25 2H, CHN), 3.1(m, 4H, PhCH₂), 2.7(m, 2H, CHCH₂), 2.41(s,
2H, OH).

Example 1AO: MS 439(M+1, 100). NMR(CDCl₃) δ 7.15-7.35(m,
10H, Ph), 4.05(s, 2H, CHOH), 3.7(m, 2H, NCH₂), 3.55(d,
30 2H, NCH), 3.0-3.2(m, 4H, PhCH₂), 2.65(s, 2H, OH),
1.78(m, 2H, NCH), 1.6(s, 2H, CH₂CH), .82(d, 12H, CHMe₂).

Example 1AH: mp 194-195 °C. MS: 467(M+1, 100). NMR(CDCl₃) δ
7.15-7.35(m, 10H, Ph), 3.95(s, 2H, CHOH), 3.65(m, 2H,
35 NCH), 3.49(d, 2H, NCH), 2.9-3.1(m, 4H, PhCH₂), 2.61(s,

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2H, OH), 2.2(m, 2H, NCH), 1.1-1.5(m, complex), .9(m, 12H, CHMe₂).

5 Example 1AM: mp 120-122 °C. MS: 495(M+1,100). NMR(CDCl₃) δ 7.1-7.35(m, 10H, Ph), 3.95(s, 2H, NCH), 3.45-3.7(m, 4H, NCH₂, NCH), 2.9-3.2(m, 4H, PhCH₂), 2.1(m, 2H, NCH₂), .78-1.45(m, complex, hexyl).

10 Example 1AN: mp 105-107 °C. MS: 523(M+1,100). NMR(CDCl₃) δ 7.18-7.31(m, 10H, Ph), 3.95(s, 2H, CHOH), 3.7(m, 2H, NCH₂), 3.56(d, 2H, NCH), 2.9-3.15(m, 4H, PhCH₂), 2.18(m, 2H, NCH₂), .9-1.45(m, complex, hexyl).

15 Example 1AL: mp 144-145 °C. MS: 551(M+1,100). NMR(CDCl₃) δ 7.15-7.35(m, 10H, Ph), 3.95(s, 2H, CHOH), 3.65(m, 2H, NCH₂), 3.55(d, 2H, NCH), 2.9-3.15(m, 6H, PhCH₂, OH), 2.15(m, 2H, NCH₂), .9-1.5(m, complex, heptyl).

20 Example 1AC: mp 160-163 °C. MS: 463(M+1,100). NMR(CDCl₃) δ 7.15-7.35(m, 10H, Ph), 5.06(br t, 2H, NCH₂CH), 4.1(dd, 2H, NCH₂), 3.9(s, 2H, CHOH), 3.5(d, 2H, NCH), 2.8-3.1(m, 4H, PhCH₂), 1.65(s, 6H, CHCH₃), 1.38(s, 6H, CHCH₃).

25 Example 1AJ: mp 205-207 °C. MS: 435(M+1,100). NMR(CDCl₃) δ 7.1-7.35(m, 10H, Ph), 4.85(s, 2H, C=CH), 4.58(s, 2H, C=CH), 4.28(d, 2H, NCH₂), 3.95(s, 2H, CHOH), 3.7(br d, 2H, CHOH), 2.9-3.2(m, 4H, PhCH₂), 2.5(d, 2H, NCH₂), 1.75(s, 6H, CH₃).

30 Example 1AI: MS: 509(M+1,100). NMR(CDCl₃) δ 7.0-8.4(m, 18H, Ph, pyr.), 4.8-5.0(m, 4H, NCH₂), 3.7(s, 2H, CHOH), 3.5(d, 2H, NCH), 2.9-3.2(m, 8H, PhCH₂, pyrCH₂).

35 Example 1AP: mp 198-200 °C. MS: 403(M+1,100). NMR(CDCl₃) δ 7.2-7.35(m, 10H, Ph), 4.5(s, 2H, CCH), 4.42(s, 2H, NCH),

4.1(s, 2H, CHOH), 3.8(d, 2H, NCH), 2.9-3.2(m, 4H, PhCH₂), 2.7-2.85(m, 4H, NCH₂).

5 Example 10: mp 105-106 °C. MS: 531(M+1,100). NMR(CDCl₃)δ 7.1-7.35(m, 10H, Ph), 3.0-4.15(m, complex), 2.25(m, 2H, OH).

10 Example 1S: MS: 619(M+1,100). NMR(CDCl₃)δ 7.18-7.3(m, 10H, Ph), 3.0-4.2(m, complex), 2.25(m, 2H, OH).

Example 1P: mp 80-82 °C. MS: 607(M+1,100), 257(9.6). NMR(CDCl₃)δ 7.05-7.35(m, 10H, Ph), 3.95(s, 2H, CHOH), 3.4-3.75(m, 4H, NCH, NCH₂), 2.9-3.15(m, 4H, PhCH₂), 2.1(m, 2H, OH), .85-1.6(m, complex).

15 Example 1AA: mp 70-75 °C. MS: 553(M+1,100). NMR(CDCl₃)δ 7.05-7.4(m, 10H, Ph), 3.4-4.25(m, complex), 2.9-3.15(m, 4H, PhCH₂), 2.2-2.8(m, complex).

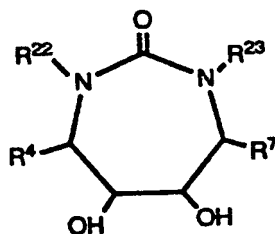
20 Example 1X: mp 210-212 °C. MS: 435(M+1,100). NMR(CDCl₃)δ 7.18-7.35(m, 10H, Ph), 4.06(s, 2H, CHOH), 3.68(br d, 2H, NCH), 3.55(q, 2H, NCHCH), 3.1(m, 4H, PhCH₂), 2.55(s, 2H, OH), 2.05(q, 2H, CHCH), .9(m, 2H, NCH₂CH), .42(m, 4H, -CH₂-), .008(m, 4H, -CH₂-).

<u>Example</u>		
<u>Number</u>		<u>MS (M+1)</u>
5	1B	327.9
	1D	406.5
	1H	407
	1J	531
	1K	463
10	1L	604.5
	1M	619
	1N	607
	1O	467
	1R	473
15	1T	604
	1U	793
	1V	411
	1AE	395
	1AG	355
20	1AS	563
	1AT	435
	1BD	409
	1AQ	507.26
	1AU	559.29

- 25 Example 1AR: mp 85-87 °C.
Example 1C: mp 164-166 °C.
Example 1A: mp 170-174 °C.

30 Using the above-described techniques or variations thereon appreciated by those of skill in the art of chemical synthesis, the compounds of Tables 1b can be prepared.

Table 1b



5

Example Number	R22	R23	R4	R7
1BE	methyl	phenylmethyl	4-F- phenylmethyl	4-F- phenylmethyl
1BF	allyl	cyclopropyl- methyl	4-F- phenylmethyl	4-F- phenylmethyl
1BG	allyl	cyclopropyl- methyl	4-Cl- phenylmethyl	4-Cl- phenylmethyl
1BH	allyl	cyclopropyl- methyl	1-pyrrolyl- methyl	1-pyrrolyl- methyl
1BI	allyl	ethyl	4-F- phenylmethyl	4-F- phenylmethyl
1BJ	isoprenyl	isoprenyl	1-pyrrolyl- methyl	1-pyrrolyl- methyl
1BK	n-butyl	n-butyl	4-hydroxy- benzyl	4-hydroxy- benzyl
1BL	N-2- acetoamido-3- methyl-butane	cyclopropyl- methyl	benzyl	1-pyrazolyl- methyl
1BM	benzyl	cyclopropyl- methyl	benzyl	1-pyrazolyl- methyl
1BN	cyclopropyl- methyl	cyclopropyl- methyl	4-phenyl- benzyl	1-pyrrolyl- methyl

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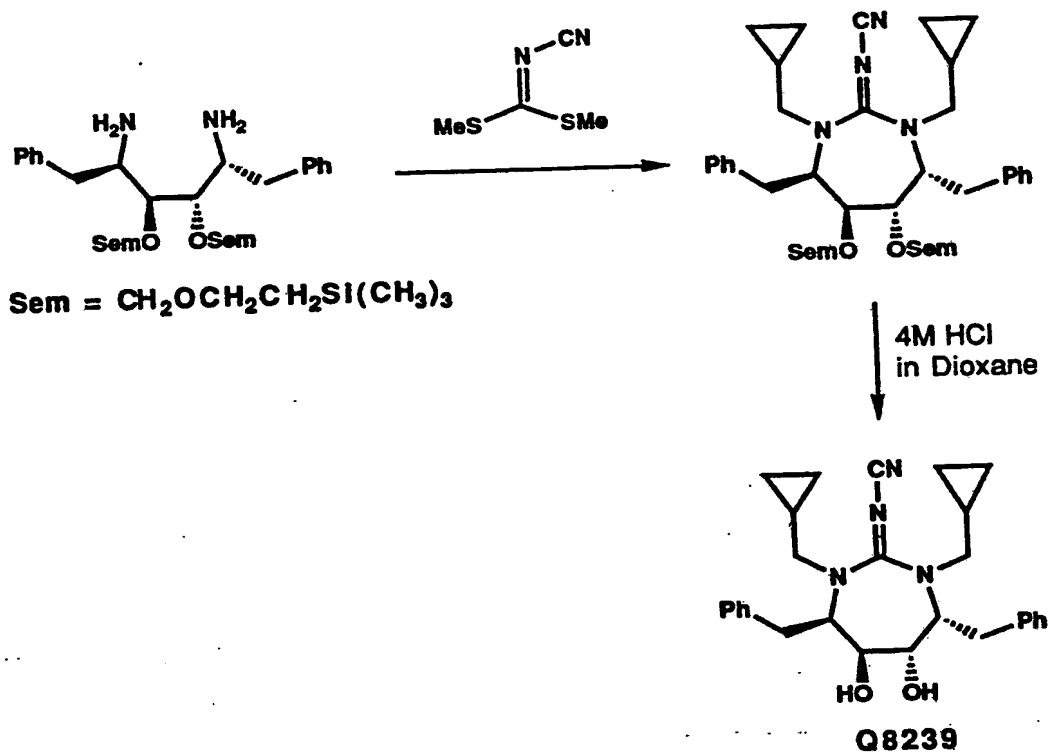
1BO	cyclopropyl- methyl	cyclopropyl- methyl	benzyl	1-imidazolyl- methyl	
1BP	cyclopropyl- methyl	cyclopropyl- methyl	benzyl	1-pyrrolyl- methyl	
1BQ	n-butyl	cyclopropyl- methyl	benzyl	1-pyrrolyl- methyl	
1BR	cyclopropyl- methyl	cyclopropyl- methyl	benzyl	1-pyrazolyl- methyl	
1BR1		cyclopropyl- methyl	1-pyrrolyl- methyl	1-pyrrolyl- methyl	
1BR2	allyl	allyl	benzyl	4-pyridinyl- methyl	
1BR3	N-butyl	cyclopropyl- methyl	benzyl	4-pyridinyl- methyl	

Synthesis of Cyclic Guanidines

5 Cyclic guanidine compounds of the invention wherein
W = NH(C=N-CN)NH, differ from the cyclic urea compounds
of the invention wherein W = NH(C=O)NH. Described below
are representative methods for the preparation of cyclic
guanidine compounds of the invention.

10

SYNTHESIS OF CYCLIC GUANIDINES



The structures of the Examples below are shown in Table 1c.

5

Example 1BS (08239)

SYNTHESIS OF INTERMEDIATE A: Diamino Disem 561mg (1mmol) was dissolved in 2ml pyridine and to this solution was added 175mg (1.2mmol) Dimethyl N-cyanodithioiminocarbonate. The contents were refluxed in a 125°C oil bath for 2 hours. (Caution: Methyl mercaptan is a by-product and the reaction should be vented to a Clorox scrubber). TLC (1:2 EtOAc:Hexane Rf=0.4) indicated a complete reaction. The reaction was

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diluted with 100ml dichloromethane. The organic layer was washed with 1N HCL(2x25ml) followed by sat. sodium bicarbonate solution(25ml). It was separated and dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel(55g; using 1:3 followed by 1:2 EtOAc:Hexane) to provide 372mg(60.9% yield) of the desired intermediate A as a colorless oil.

10 SYNTHESIS OF INTERMEDIATE B: Intermediate A 305mg(0.5mmol) was dissolved in 2ml dimethylformamide and to this solution, cooled in a 0°C ice bath, was added NaH(60% in oil) 80mg(2mmol) slowly. The contents were stirred at room temperature for 30 minutes. The
15 mixture was cooled in a 0°C ice bath and (bromomethyl)cyclopropane 0.19ml(2mmol) was added via syringe and stirred at room temperature for 18 hours. TLC(1:4 EtOAc:Hexane R_f=0.31) indicated a complete
20 reaction. The reaction was worked up by diluting with water(50ml) and extracting with diethylether (2x25ml). The organic layer was dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel(33g; 1:5 followed by 1:4 EtOAc:Hexane) to provide 243mg(67.6% yield) of the desired intermediate B
25 as a colorless oil.

SYNTHESIS OF EXAMPLE 1BS: Intermediate B 110mg (0.153mmol) was placed in a 10ml R.B. Flask and cooled in a 0°C ice bath. To the flask was added 4M HCl in
30 dioxane 1ml (4mmol) and the mixture stirred at room temperature for 15 minutes. TLC(1:1 EtOAc:Hexane R_f=0.15) indicated a complete reaction. The mixture was worked up by quenching in sat. sodium bicarbonate(25ml) and extracting with dichloromethane (2x25ml). The
35 organic extracts were dried over magnesium sulfate and the filtrate taken to dryness. The residue was

purified on SiO₂ gel(33g; 1:1 EtOAc:Hexane followed by 10:1:10 EtOAc:EtOH:Hexane) to provide 27mg(38.5% yield) of the desired Q8239 as a white solid. m.p.211.2°C

Example 1BT

5

SYNTHESIS OF INTERMEDIATE C: Intermediate A 1.515g(2.48mmol) was dissolved in 7.5ml dimethylformamide and to this solution, cooled in a 0°C ice bath, was added NaH(60% in oil) 397mg(9.92mmol) slowly. The contents were stirred at room temperature for 30 minutes. The mixture was cooled in a 0°C ice bath and p-benzyloxybenzyl chloride 2.308g(9.92mmol) was added as a solid and the mixture stirred at room temperature for 18 hours. TLC(1:4 EtOAc:Hexane R_f=0.31) indicated a complete reaction. The reaction was worked up by diluting with water(100ml) and extracting with diethylether (2x50ml). The organic layer was dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel(130g; 1:4 followed by 1:3 EtOAc:Hexane)to provide 2.068g(83.1% yield) of the desired intermediate C as a colorless foam.

SYNTHESIS OF EXAMPLE 1BT: Intermediate C 1.928g (1.92mmol) was placed in a 100ml R.B. flask and cooled in a 0°C ice bath. To the flask was added 4M HCl in dioxane 15ml (60mmol) and the mixture stirred at room temperature for 15 minutes. TLC(1:1 EtOAc:Hexane R_f=0.25) indicated a complete reaction. The mixture was worked up by quenching in 0.5N sodium hydroxide solution(100ml) and washing with sat.sodium bicarbonate(50ml) and extracting with dichloromethane (3x50ml). The organic extracts were dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel(130g; 1:1

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EtOAc:Hexane) to provide 1.284g(89.9% yield) of the desired Q8241 as a white solid. m.p.90.1°C

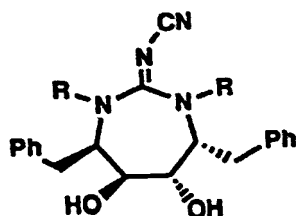
Example 1BU

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SYNTHESIS OF EXAMPLE 1BU: Example Number 1BT

1.161g(1.56mmol) was dissolved in 15ml of ethanol. To the mixture was added 1.1g of 5% Palladium on Carbon and the suspension stirred for 18 hours under hydrogen(1 atm). TLC indicated an incomplete reaction. The mixture was treated with 1.1g of 10% Palladium Hydroxide on Carbon and stirred for 2 hours under hydrogen(1 atm). TLC(10:1:10 EtOAc:Hexane Rf=0.31) indicated a complete reaction. The suspension was filtered through a celite pad and the filtrate taken to dryness. The residue was purified on SiO₂ gel(130g; 10:1:10 followed by 10:2:10 EtOAc:EtOH:Hexane) to provide 458mg(52.2% yield) of the desired Q8242 as a white solid. m.p.103.3°C

Table 1c



Example Number	R	Ki	IC90	M+H (M+NH ₄)	m.p. (°C)
1BS	cyclopropylmethyl	++	+++	459	211.2
1BT	CH ₂ (C ₆ H ₄) -p-OCH ₂ C ₆ H ₅	++	+	743	90.1
1BU	CH ₂ (C ₆ H ₄) -p-OH	+++	+++	563	103.3
1BV	cyclopentylmethyl	+++	+++	515	99.7
1BW	allyl	++	++	431	70.2
1BX	n-butyl	+++	+++	463	58.3
1BY	Beta-naphthylmethyl	++	+	631	111.1
1BZ	benzyl	++	+++	531	94.0
1CA	CH ₂ (C ₆ H ₄) -m-OCH ₂ C ₆ H ₅	++	+	743	75.2
1CB	p-nitrobenzyl	++	++	621	130.3
1CC	m-nitrobenzyl				114
1CD	CH ₂ (C ₆ H ₄) -m-OH	+++	+++	563	124

1CE	p-aminobenzyl	++	+++	561	226.3
1CF	m-aminobenzyl	+++	+++	561	114.4
1CG	p-nitrilobenzyl	++	+++	581	
1CH	m-nitrilobenzyl	++	+++	(598)	108.5
1CI	dimethylallyl	++	+++	487	72.9
1CJ	cyclohexylmethyl	+++	+++	543	96.4
1CK	cyclobutylmethyl	+++	+++	487	235.5
1CL	propyl	++	+++	435	216.6
1CM	3-methyl-1-butyl	++	+++	491	76.1
1CN	CH ₂ (C ₆ H ₄)-p-CH ₂ OH	++	+++	591	100.9
1CO	CH ₂ (C ₆ H ₄)-m-CH ₂ OH	+++	+++	591	193.8
1CP	CH ₂ (C ₆ H ₄)-m-CHO	+++	+++	587	
1CQ	CH ₂ (C ₆ H ₄)-m-CH=N-OH	+++	+++	617	121.2

The structures of the Examples below are shown in Tables 1d and 1e.

5

Example 7F

A solution of Example 1X (120 mg, 0.27 mmol) in methylene chloride (25 mL) was cooled in an ice bath at 0 °C and treated with triethylamine (110 mg, 1.1 mmol).

Then a solution of thionyl chloride (150 mg, 1.3 mmol) in methylene chloride (10 mL) was added dropwise. The mixture was stirred for 10 minutes and then washed with sat'd $\text{NaHCO}_3(\text{aq})$, brine, and dried over MgSO_4 . The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (50% EtOAc/Hexanes) to give 100 mg of a white foam. HRMS calculated for $\text{C}_{27}\text{H}_{32}\text{N}_2\text{O}_4\text{S}$: 481.2161; found: 481.2152.

10

Example 7G

A solution of Example 1X (120 mg, 0.27 mmol) in methylene chloride (25 mL) was cooled in an ice bath at 0 °C and treated with triethylamine (80 mg, 0.8 mmol). Then a solution of trichloromethyl chloroformate (DIPHOSGENE) (53 mg, 0.27 mmol) in methylene chloride (5 mL) was added dropwise. The mixture was stirred for 60 minutes and then washed with sat'd $\text{NaHCO}_3(\text{aq})$, brine, and dried over MgSO_4 . The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is chromatographed on silica gel (50% EtOAc/Hexanes) to give 90 mg of a white foam. MS NH_3 $(\text{M}+\text{H})^+ = 461$.

25

Example 7I

A solution of Ex. 1X (500 mg, 1.15 mmol) in THF (25 mL) was treated with thiocarbonyldiimidazole (410 mg, 2.3 mmol) and heated to reflux for 1.5 hours. The solvent is removed on a rotary evaporator and the resulting residue is chromatographed on silica gel (50% EtOAc/Hexanes) to give 320 mg of a white solid. MS NH_3 $(\text{M}+\text{H})^+ = 477.3$

35

Example 7H and 7U

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- A solution of Example 7I (220 mg, 0.46 mmol) in dry toluene (25 mL) was heated to reflux and treated with Bu_3SnH (0.4 mL, 1.5 mL) and 20 mg of AIBN. The mixture was heated at reflux for 2.5 hours. The solvent is removed on a rotary evaporator and the resulting residue is chromatographed on silica gel (20% EtOAc/Hexanes) to give 70 mg of 7H as a white solid. MS NH_3 $(\text{M}+\text{H})^+ = 447.2$.
- A later fraction gave 30 mg of Example 7U as a white solid. MS $(\text{M}+\text{H})^+ 419.2$.

Example 7V

- A solution of Ex. 1X (112 mg, 0.26 mmol) in methylene chloride (5 mL) was cooled in an ice bath at 0 °C and treated with DAST (110 mg, 1.1 mmol). The mixture was stirred for 10 minutes and then washed with sat'd $\text{NaHCO}_3(\text{aq})$, brine, and dried over MgSO_4 . The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (65% EtOAc/Hexanes) to give 40 mg of a colorless residue. HRMS calculated for $\text{C}_{27}\text{H}_{33}\text{N}_2\text{O}_2\text{F}$: 437.2604; found: 437.2593.

25

Example 7o

- A solution of Ex. 3U (600 mg, 1.9 mmol) in pyridine (10 mL) is treated with methanesulfonyl chloride (170 mg, 1.5 mmol) and stirred at room temperature for 3 hours. The mixture is quenched with 5 mL of methanol. The solution is concentrated on a rotary evaporator and the resulting residue is dissolved in ethyl acetate and then washed with dilute $\text{HCl}(\text{aq})$, brine, and dried over MgSO_4 . The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is

chromatographed on silica gel (40% EtOAc/Hexanes) to give Example 7o (420 mg) as a white solid. ms (M+H)⁺ = 685.

5

Example 7Y

A solution of Example 7o (100 mg, 0.15 mmol) in DMF (4 mL) was treated with NaN₃ (100 mg, 1.5 mmol) and heated at 80 °C for 2 hrs and then stirred overnight at 40 °C. The solution was diluted with water and the resulting white solid is extracted into ethyl acetate. The organic extract is washed with water, brine, and dried over MgSO₄. The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (50% EtOAc/Hexanes) to give Example 7Y (80 mg) as a white solid. MS (M+H)⁺ = 632. IR (CHCl₃) 2214 cm⁻¹ for N₃.

20

Example 7J and 7K

A solution of Ex. 3U (100 mg, 0.165 mmol) in pyridine (3 mL) is treated with acetic anhydride (84 mg, 0.824 mmol) and stirred at room temperature for 2 hours. The mixture is quenched with 5 mL of methanol. The solution is concentrated on a rotary evaporator and the resulting residue is dissolved in ethyl acetate and then washed with dilute HCl(aq), brine, and dried over MgSO₄. The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (40% EtOAc/Hexanes) to give Example 7J (24 mg) as a white solid. HRMS: (M+H)⁺ calculated for C₄₅H₄₃N₂O₅: 691.317198; found: 691.316252.

30

Also gives as a later fraction Example 7K (27 mg) as a white solid. HRMS: (M+H)⁺ calculated for C₄₃H₄₁N₂O₄: 649.306633; found: 649.304918.

35

Example 7R

5 A solution of Ex. 3U (100 mg, 0.165 mmol) in methylene chloride (5 mL) is treated with 2,2dimethoxy propane (174 mg, 1.65 mmol), p-toluenesulfonic acid (10 mg) and stirred at room temperature for overnight. The mixture is diluted with methylene chloride and washed with saturated NaHCO_3 , brine, and dried over MgSO_4 . The
10 solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (50% EtOAc/Hexanes) to give Example 7R (73 mg) as a white solid. HRMS: $(\text{M}+\text{H})^+$ calculated for $\text{C}_{44}\text{H}_{43}\text{N}_2\text{O}_3$: 647.327369; found: 647.327531

15

Example 7P

A solution of Ex. 3U (200 mg, 0.33mmol) in methylene chloride (5 mL) is treated with oxalyl
20 chloride (249 mg, 2.0 mmol) and stirred at room temperature for overnight. The solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (50% EtOAc/Hexanes) to give a tan solid. This was recrystallized from
25 CH_2Cl_2 /hexane to give Example 7P (80 mg) as a white solid. MS: $(\text{M}+\text{H})^+$ 661.4.

Example 7S

30 A solution of Ex. 3U (100 mg, 0.165 mmol) in methylene chloride (5 mL) is treated with trimethylorthobutyrate (244 mg, 1.65 mmol) p-toluenesulfonic acid (10 mg) and stirred at room temperature for 30 minutes. The mixture is diluted with
35 methylene chloride and washed with saturated NaHCO_3 , brine, and dried over MgSO_4 . The solution is filtered

and the solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (50% EtOAc/Hexanes) to give Ex. 7S. MS: (M+H)⁺ 691.5

5

Example 7B

A solution of the Di-MEM protected Ex. 1C (550 mg, 0.94 mmol) in pyridine (11 mL) is treated with P₂S₅ (420 mg, 0.94 mmol) and heated to reflux for 3 hrs. The pyridine is evaporated off on a rotary evaporator and the residue is taken up in methylene chloride and washed with water, NaHCO₃, and brine. The solution is dried over MgSO₄, filtered and the solvent removed on a rotary evaporator and the resulting residue is chromatographed on silica gel (40% EtOAc/Hexanes) to give Example 7B (120 mg) as a clear oil. HRMS: (M+H)⁺ calculated for C₂₇H₃₃N₂O₃S: 465.221190; found: 465.220899.

20

Example 7A

A solution of Example 1C (330 mg, 0.8 mmol) in pyridine (5 mL) is treated with acetic anhydride (160 mg, 1.6 mmol) and stirred at room temperature for 4 hours. The mixture is quenched with 5 mL of methanol. The solution is concentrated on a rotary evaporator and the resulting residue is dissolved in ethyl acetate and then washed with dilute HCl (aq), brine, and dried over MgSO₄. The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is chromatographed on silica gel (50% EtOAc/Hex) to give Example 7A (120 mg) as a white solid. MS: (CI, NH₃) (M+H)⁺ = 449.1.

35

Example 7W

A solution of example 1C (160 mg, 0.39 mmol) in methylene chloride (5 mL) was cooled in an ice bath at 0 °C and treated with DAST (63 mg, 0.4 mmol). The mixture was stirred for 10 minutes and then washed with sat'd NaHCO₃(aq), brine, and dried over MgSO₄. The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (50% EtOAc/Hex) to give 80 mg of a colorless residue. HRMS calculated for C₂₅H₃₀N₂O₂F: 409.2291; found: 409.2291.

Example 7X

A solution of example 3U (100 mg, 0.16 mmol) in methylene chloride (5 mL) was cooled in an ice bath at 0 °C and treated with DAST (26 mg, 0.16 mmol). The mixture was stirred for 10 minutes and then washed with sat'd NaHCO₃(aq), brine, and dried over MgSO₄. The solution is filtered and the solvent removed on a rotary evaporator and the resulting residue is HPLC chromatographed on silica gel (50% EtOAc/Hex) to give 35 mg of a white foam. HRMS calculated for C₄₁H₃₈N₂O₂F: 609.2917; found: 609.2911.

Example 8A

A solution of example 1X (2.06 g, 4.74 mmol) in methylene chloride was treated with diisopropylethylamine (1.53 g, 11.8 mmol), MEM-Cl (0.71 g, 5.7 mmol) and heated to reflux for 5 hours and let stir overnight at rt. The solution is concentrated on a rotary evaporator the residue is HPLC chromatographed on silica gel (5 % MeOH/CHCl₃) to give 1.3 g of the mono-MEM mono-ol intermediate. MS: (CI, NH₃) (M+H)⁺ = 523.4.

A solution of mono-MEM mono-ol intermediate from above (1.0 g, 1.9 mmol) in THF was treated with

triphenylphosphine (1.0 g, 3.8 mmol),
diethylazadicarboxylate (DEAD) (0.7 g, 4.0 mmol), and
chloroacetic acid (0.4 g, 4.2 mmol). The solution is
stirred overnight at rt. The solvent is evaporated and
5 the resulting residue is chromatographed on silica gel
(50% EtOAc/Hex) to give 0.9 g of the chloroacetate
intermediate. MS: (CI, NH₃) (M+H)⁺ = 599.3 (100%); 600
(39%).

The chloroacetate intermediate (0.9 g, 1.5 mmol)
10 was dissolved in MeOH (15 ml) and treated with NaOH(aq)
(4 ml, 1N) and stirred at rt for 15 min. The solution
was evaporated to dryness and the residue partitioned
between water and ethyl acetate. The organic layer was
washed with water and brine and then dried over MgSO₄.
15 The solution is filtered, concentrated and the
residue is HPLC chromatographed on silica gel (85%
EtOAc/Hex) to give 400 mg of example 8A. MS: (CI, NH₃)
(M+H)⁺ = 523.4.

20

Example 7Z

A solution of example 8A (100 mg, 0.2 mmol) in MeOH
is cooled in an ice bath and treated with HCl (g) for 20
min and then stirred for an additional 40 min at 0°C.
25 The solution is then evaporated to dryness at rt and the
residue is HPLC chromatographed on silica gel (80%
EtOAc/Hex) to give 48 mg of example 7Z as a white foam.
MS: (CI, NH₃) (M+H)⁺ = 435.2

30

Example 8C

A solution of example 8A (160 mg, 0.3 mmol) in
methylene chloride (10 ml) is cooled to 0°C in an ice
bath and treated with DAST (50 mg, 0.3 mmol). The
35 solution is stirred at rt for 15 min and then quenched
with water. The organic layer is washed with water and

brine and dried over MgSO_4 . The solution is filtered, the solvent evaporated and the residue HPLC chromatographed on silica gel (50% EtOAc/Hex) to give 100 mg of example 8C. MS: $(\text{CI}, \text{NH}_3) (\text{M}+\text{H})^+ = 525.4$.

5

Example 8B

A solution of example 8C (70 mg, 0.13 mmol) in MeOH is cooled in an ice bath and treated with HCl (g) for 20 min and then stirred for an additional 40 min at 0°C . The solution is then evaporated to dryness at rt and the residue is HPLC chromatographed on silica gel (80% EtOAc/Hex) to give 40 mg of example 8B as a white foam. MS: $(\text{CI}, \text{NH}_3) (\text{M}+\text{H})^+ = 437.3$

15

Example 7AA

To a stirred suspension of 750 mg (1.72 mmol) of the diol (1X) in 35 mL of methylene chloride was added 445 mg (3.45 mmol) of diisopropylethylamine and 322 mg (2.59 mmol) of MEM chloride. After stirring 5 days the resulting solution was washed with dilute HCl, brine and was dried with anhydrous MgSO_4 . The solvent was removed under reduced pressure and the residue was chromatographed on silica gel. Elution with 50% ethyl acetate in hexanes gave 430 mg (48%) of the mono protected ether (XXVa). MS: 523 ($\text{M}+1, 100$); NMR (CDCl_3): δ 7.20 (m, 10H), 4.96 (s, 2H), 4.08 (m, 1H), 3.90 (m, 2H), 3.61 (m, 7H), 3.42 (s, 3H), 3.13 (m, 4H), 1.99 (m, 2H), 0.88 (m, 2H), 0.40 (m, 4H), 0.06 (m, 4H),

30

To a stirred solution of 78mg (0.15mmol) of Compound (XXVa) in 3 ml of methylene chloride was added 60 mg (0.74 mmol) of sodium acetate and 95 mg (0.44 mmol) of PCC. The resulting suspension was stirred 3 days and was diluted with ether and filtered through

35

florisil. The solvent was removed under reduced pressure and the residue was chromatographed on silica gel. Elution with 2.5% methanol in methylene chloride gave 68 mg (885) of example 7AA. MS:521 (m+1,100); NMR (CDCl₃): δ 7.21 (m,10H), 4.90 (d,1H), 4.70 (dd,2H), 4.10 (t,1H), 3.80-3.37 (m,8H), 3.36 (s,3H), 3.26-2.82 (m,4H), 2.22 (q,1H), 1.03 (m,2H), 0.51 (m,4H), 0.20 (m,4H),

10 Oxidation of monoprotected diol: Preparation of (XXVb):

To a stirred solution of 51 mg (0.10 mmol) of example 7AA in 4 mL of methanol was added 1 mL of concentrated HCl. The resulting solution was stirred 5h and the product was precipitated by adding water. The suspension was extracted with methylene chloride and the combined organic layers were dried with anhydrous MgSO₄. The solvent was removed under reduced pressure and the residue was chromatographed on silica gel. Elution with 33% ethyl acetate in hexanes gave 34 mg (80%) of the ketol (XXVb). MS:433 (m+1,100); NMR (CDCl₃): δ 7.21 (m,10H), 4.82 (m,1H), 4.24 (t,1H), 3.85 (m,1H), 3.74 (d,1H), 3.44 (m,3H), 3.22-2.73 (m,4H), 2.27 (q,1H), 1.01 (m,2H), 0.51 (m,4H), 0.20 (m,4H),

25

Example 7AC

To a stirred solution of 37 mg (0.09 mmol) of the ketol (XXVb) in 4 mL of ethanol and 2 mL of water was added 40 mg (0.48 mmol) of methoxylamine hydrochloride. The resulting solution was stirred overnight and the product was precipitated by adding water. The suspension was extracted with methylene chloride and the combined organic layers were dried with anhydrous MgSO₄. The solvent was removed under reduced pressure to give 40 mg (100%) of example 7AC. MS: 462 (m+1,100); NMR

35

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(CDCl₃) δ 7.20 (m, 10H), 5.30 (m, 2H), 4.80 (t, 1H), 4.24 (t, 1H), 3.77 (m, 4H), 3.40 (m, 3H), 2.90 (m, 4H), 2.25 (dd, 1H), 1.03 (m, 2H), 0.48 (m, 4H), 0.23 (m, 2H), 0.12 (m, 2H).

5

Example 7AB

By substituting hydroxylamine hydrochloride in the above procedure, the desired product can be obtained:
 10 MS: 448 (m+1, 100); NMR (CDCl₃): δ 8.13 (s, 1H), 7.20 (m, 10H), 5.42 (t, 1H), 4.83 (t, 1H), 3.75 (m, 2H), 3.40 (m, 3H), 2.94 (m, 4H), 2.22 (dd, 1H), 1.17 (m, 1H), 0.90 (m, 1H), 0.47 (m, 4H), 0.23 (m, 2H), 0.11 (m, 2H).

15

Example 7AD

To a stirred solution of 98 mg (0.23 mmol) of ketol (XXVb) in 5 mL of butanol was added 57 mg (0.71 mmol) of formamidine hydrochloride and 40 mg (0.75 mmol) of sodium methoxide. The resulting suspension was stirred
 20 30 min and was refluxed overnight. The butanol was removed under reduced pressure and water was added. The suspension was extracted with methylene chloride and the combined organic layers were dried with anhydrous MgSO₄.
 25 The solvent was removed under reduced pressure and the residue was chromatographed on silica gel. Elution with 25% ethyl acetate in hexanes gave 28 mg (28%) of example 7AD. MS: 442 (m+1, 100); NMR (CDCl₃): δ 7.58 (s, 1H), 7.20 (m, 10H), 4.55 (m, 2H), 3.51 (m, 2H), 3.24 (m, 2H), 3.05 (m, 2H), 2.53 (q, 1H), 2.27 (q, 1H), 0.91 (m, 2H), 0.45 (m, 4H), 0.11 (m, 4H).
 30

Examples 7AE and 7AF

35 To a stirred solution of 65 mg (0.15 mmol) of the diol (1X) in 1 mL of DMF was added 5.5 mg (0.18 mmol) of

80% sodium hydride. The resulting suspension was stirred 20 min. and 68 mg (0.48 mmol) of methyl iodide was added. After stirring overnight, the suspension was quenched with water, extracted with ethyl acetate, and the combined organic layers were dried with anhydrous MgSO₄. The solvent was removed under reduced pressure and the residue was chromatographed on silica gel. Elution with 25% ethyl acetate in hexanes gave 19 mg (28%) of example 7AE along with 25 mg (37%) of Example 7AF.

Example 7AE: MS: 449 (m+1,100); NMR (CDCl₃): δ 7.26 (m,10H), 4.05 (dd,1H), 3.84 (m,1H), 3.67 (m,2H), 3.60 (s,3H), 3.53 (m,2H), 3.13 (m,3H), 2.90 (m,2H), 1.94 (dt,2H), 0.89 (m,2H), 0.41 (m,4H), 0.04 (m,4H),

Example 7AF: MS: 463 (m+1,100); NMR (CDCl₃): δ 7.21 (m,10H), 3.67 (m,4H), 3.62 (s,6H), 3.58 (s,2H), 3.10 (m,4H), 1.92 (dd,2H), 0.83 (m,2H), 0.41 (m,4H), 0.04 (m,4H).

Examples 7AG and 7AH

By substituting benzyloxymethyl chloride in the above procedure, example 7AG and 7AH were obtained.

Example 7AG: MS: 615 (m+1,100); ¹H NMR (CDCl₃): δ 7.50-7.06 (m,20H), 4.88 (ab,4H), 3.92 (s,2H), 3.56 (m,2H), 3.47 (dd,2H), 3.14 (m,4H), 1.88 (dd,2H), 0.62 (m,2H), 0.34 (m,4H), 0.05 (m,4H).

Example 7AH: MS: 525 (m+1,100); ¹H NMR (CDCl₃): δ 7.45-7.10 (m,15H), 4.74 (ab,2H), 4.13 (dd,1H), 3.82-3.50 (m,5H), 3.09 (m,4H), 2.76 (s,1H), 1.95 (dt,2H), 0.92 (m,2H), 0.40 (m,4H), 0.03 (m,4H).

Examples 7AI and 7AJ

By substituting allyl bromide in the above procedure, example 7AI and 7AJ were obtained.

5 Example 7AI: ^1H NMR (CDCl_3) δ 7.26 (m, 10H), 6.05 (m, 2H), 5.30 (dd, 4H), 4.28 (m, 2H), 3.76 (s, 2H), 3.60 (m, 4H), 3.10 (m, 4H), 1.93 (dd, 2H), 0.86 (m, 2H), 0.40 (m, 4H), 0.01 (m, 4H).

10 Example 7AJ: MS: 475 (m+1, 100); ^1H NMR (CDCl_3) δ 7.27 (m, 10H), 6.01 (m, 1H), 5.32 (dd, 2H), 4.34 (dd, 1H), 4.18 (dd, 1H), 4.66 (m, 5H), 3.10 (m, 4H), 2.82 (s, 1H), 1.95 (m, 2H), 0.85 (m, 2H), 0.40 (m, 4H), 0.04 (m, 4H).

15

Example 8E

A solution of Example 5F (500 mg, 0.7 mmol) in methylene chloride (10 ml) is cooled to 0°C in an ice bath and treated with DAST (112 mg, 0.7 mmol). The solution is stirred at 0°C for 15 min and then quenched with sat'd NaHCO_3 . The organic layer is washed with water and brine and dried over MgSO_4 . The solution is filtered, the solvent evaporated and the residue HPLC chromatographed on silica gel (50% EtOAc/Hex) to give 250 mg of example 8E as a white foam. MS: (CI, NH_3) $(\text{M}+\text{H})^+ = 721$

30

Example 8F

A solution of example 8E (200 mg, 0.28 mmol) in MeOH is cooled in an ice bath and treated with gaseous HCl for 20 min and then stirred for an additional 40 min at 0°C . The solution is then evaporated to dryness at rt and the residue is HPLC chromatographed on silica gel

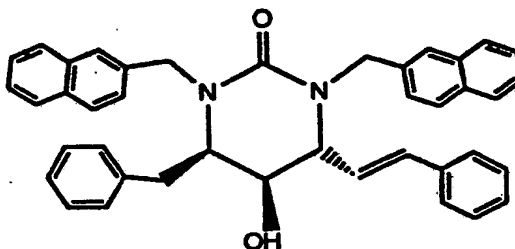
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(5% MeOH/CHCl₃) to give 120 mg of example 8F as a white foam. MS: (CI, NH₃) (M+H)⁺ = 541

Example 8G (via Alkene Intermediate XXIX):

5 A solution of Example 7Q (150 mg, 0.22 mmol) in DMF was treated with sodium iodide (160 mg, 1.1 mmol) and heated at 90°C for 2 hrs. The mixture is cooled to room temperature, diluted with water and the precipitate is
10 extracted into CH₂Cl₂. The extract is washed with water and brine, dried over MgSO₄ and evaporated to give a yellow oil. This is HPLC chromatographed on silica gel (50% EtOAc/Hex) to give 50 mg of alkene intermediate (XXIX) as a white solid. MS: (CI, NH₃) (M+H)⁺ = 589

15



Alkene intermediate (XXIX)

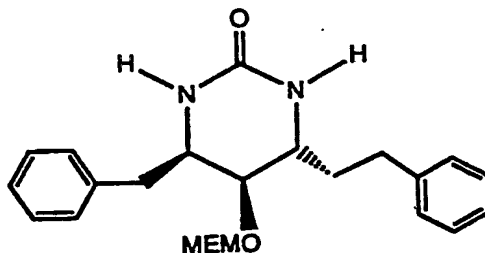
20 A solution of alkene intermediate (XXIX) (40 mg, 0.07 mmol) in THF was treated with 20 mg of 10% Pd/C and hydrogenated in a Parr Hydrogenator at 50 psi overnight. The catalyst was filtered off and the filtrate concentrated. The resulting residue was HPLC
25 chromatographed on silica gel (70% EtOAc/Hex) to give 10 mg of Example 8G as a white solid. MS: (CI, NH₃) (M+H)⁺ = 591.5

Example 8H

30

Method 1.

A. Synthesis of 6-membered ring cyclic urea (XXX):



(XXX)

5 The synthesis of the six-membered cyclic urea (XXX) is outlined in Scheme 8. A solution of N-Cbz-D-phenylalanine N,O-dimethylhydroxylamide (33.5 g, 0.098 mol) in ether was cooled to 0°C and treated with 300 mL of a 1 M solution of vinyl magnesium bromide in THF. The mixture was stirred for 30 mins and then poured into an ice cold solution of 1 N HCl (500 mL). The mixture was extracted into ether and the extracts washed with water and brine. The organic layer was dried over MgSO₄, filtered and concentrated to give the desired vinyl ketone as a thick, light yellow residue which was used without further purification. MS: (CI, NH₃) (M+H)⁺ = 310 (77%); (M+NH₄)⁺ = 327.1 (100%).

15 The crude ketone was dissolved in methanol (350 mL) and treated with cerium trichloride heptahydrate (37.2 g, 0.1 mol) and cooled in an ice bath. While stirring vigorously sodium borohydride (3.78 g 0.1 mol) was added slowly, a small portion at a time, over a period of 30 min. After the addition was complete the mixture was allowed to warm to room temperature and stirred for an additional 1 hr. The solvent was removed under vacuum on a rotary evaporator and the residue was partitioned between 1 N HCl and methylene chloride. The organic layer was washed with water, brine, and then dried over MgSO₄, filtered and concentrated to give the desired

allylic alcohol as an off-white solid which was used without further purification.

A solution of the crude allylic alcohol and diisopropylethylamine (30 g, 0.23 mol) in methylene chloride was cooled in an ice bath and treated dropwise with methanesulfonyl chloride (28 g, 0.24 mol). The solution was stirred for 30 mins, then washed sequentially with 1 N HCl, water, brine and dried over MgSO_4 . The solution was filtered and concentrated to give the crude mesylate as a thick oil. To a flamed-dried flask was added copper cyanide (12 g, 0.144 mol) and 100 mL of THF. The flask was cooled to -78°C under nitrogen atmosphere. A solution of benzylmagnesium chloride (360 mL, 2M in THF, 0.72 mol) was added via syringe and the resulting thick solution was stirred at -60°C for 20 mins and at 0°C for 30 mins. The solution was then cooled to -78°C and a solution of the mesylate in 130 mL of THF was added via syringe. The solution was stirred at -60°C for 45 mins and then poured into a mixture of 1 N HCl/ice. This was extracted into ethyl acetate and the organic layer was washed sequentially with NH_4Cl (aq), NH_4OH , brine, dried over MgSO_4 , filtered and concentrated. The resulting residue is chromatographed on silica gel (hexane, then 10% EtOAc/Hex) to give 11.7 g of the desire alkene as a white solid. MS: (CI, NH_3) $(\text{M}+\text{H})^+ = 386.3$ (98%); $(\text{M}+\text{NH}_4)^+ = 403.2$ (100%).

A solution of the above alkene (11.0 g, 0.029 mol) in methylene chloride (75 mL) was cooled to 0°C in an ice bath and treated with 60% m-chloroperbenzoic acid (14.0 g, 0.049 mol). The solution was stirred 0°C for 7 hrs until TLC analysis showed no starting material remained. A precipitate formed during this time. The suspension was diluted with methylene chloride and washed sequentially with 1 N $\text{Na}_2\text{S}_2\text{O}_3$, 1 N sodium hydroxide, water, brine, dried over MgSO_4 , filtered and

concentrated to give the epoxide as a thick oil which was used without further purification.

To solution of crude epoxide in 80 mL of DMF was added sodium azide (20 g , 0.3 mol), ammonium chloride (2.5 g , 0.047 mol) and 20 mL of water. The mixture was heated at 90 °C for 3 hrs and then stirred at rt overnight. The solvent was removed under high vacuum on a rotary evaporator and the residue was partitioned between water and methylene chloride. the organic layer was washed with water and brine, dried over MgSO_4 , filtered and concentrated to give a residue. This was then chromatographed on silica gel (20% EtOAc/Hex) to give 7.4 g of the azide alcohol as a white solid. MS: $(\text{Cl}, \text{NH}_3) (\text{M}+\text{H})^+ = 445.0$ (25%); $(\text{M}+\text{NH}_4-\text{BnOH})^+ = 462.2$ (100%).

A solution of the azide alcohol above (7.2 g, 0.016 mol) in methylene chloride was treated with diisopropylethylamine (4.2 g , 0.032 mol) and MEM-Cl (4.0 g 0.032 mol) and heated to reflux overnight (18 hrs). The mixture was concentrated and the residue chromatographed on silica gel (20% EtOAc/Hex - 35% EtOAc/Hex) to give 7.7 g of the MEM protected azido alcohol as a colorless oil. MS: $(\text{Cl}, \text{NH}_3) (\text{M}+\text{H})^+ = 533.2$ (100%).

To a solution of MEM protected azido alcohol (5.7 g 0.0107 mol) in ethyl acetate was added 2 mL of acetic acid and 1 g of Pearlman's catalyst (10 % $\text{Pd}(\text{OH})_2$ on Carbon) and the solution was hydrogenated at 55 psi for 22 hrs. The solution was filtered through Celite and the filtrate was extracted with 1 N HCl (organic layer turn orange). The acidic aqueous extract was made basic with 50% NaOH (while cooling in an ice bath) and the precipitate is extracted into ethyl acetate. The organic layer is washed with water, brine, dried over MgSO_4 , filtered and concentrated to give 2.5 g of the MEM

protected diamino alcohol as a colorless oil. MS: (CI, NH₃) (M+H)⁺ = 373.1 (100%).

To a solution of the MEM protected diamino alcohol (2.5 g 0.0067 mol) in THF was added 1,1-carbonyldiimidazole (1.1 g, 0.0067 mol) and stirred over night at room temperature. The solution was concentrated and the residue partitioned between 1 N HCl and CH₂Cl₂. The organic layer is washed with brine, dried over MgSO₄, filtered and concentrated. The residue is HPLC chromatographed on silica gel (5% MeOH/CHCl₃) to give 1.2 g of the MEM protected 6-membered ring cyclic urea (XXX) as a white solid. MS: (CI, NH₃) (M+H)⁺ = 399.1 (100%).

B. The MEM protected 6-membered ring cyclic urea (XXX) (100 mg, 0.27 mmol) was alkylated with cyclopropylmethylbromide (250 mg , 1.8 mmol) followed by removal of the MEM group, as described in general procedure 5, to give after chromatography on HPLC (silica gel, 10% MeOH/CHCl₃) 20 mg of Example 8H as a clear, viscous residue. MS: (CI, NH₃) (M+H)⁺ = 419.4 (100%).

Method 2.

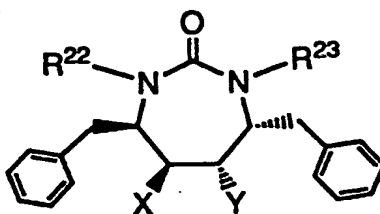
A solution of Example 8A (160 mg, 0.3 mmol) and thiocarbonyldiimidazole (55 mg, 0.3 mmol) in THF was heated to reflux for 4 hrs. The mixture was evaporated and the residue chromatographed on silica gel (50% EtOAc/Hex) to give 34 mg (0.055 mmol) of the corresponding thiocarbamate. The thiocarbamate was dissolved in 2 mL of toluene and heated to reflux. To the refluxing solution was added tributyltin hydride (32 mg, 0.1 mmol) and 2 mg of AIBN. The mixture was refluxed for 1 hour, concentrated, and the residue chromatography on HPLC (silica gel, 65% EtOAc/Hex) to give 20 mg of clear colorless oil. The oil was dissolved in MeOH, cooled in an ice bath and gaseous HCl was bubbled

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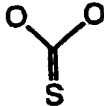
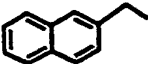
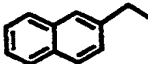
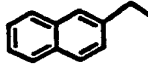

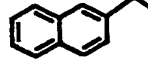
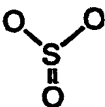
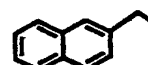
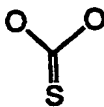
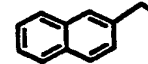
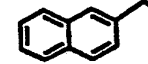
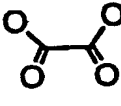
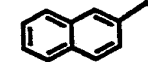
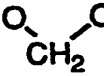

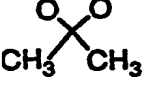
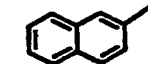
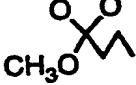
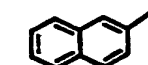
through the solution for 30 mins. The solution was then stirred at room temperature overnight, concentrated and the residue chromatography on HPLC (silica gel, 10% MeOH/CHCl₃) to give 10 mg of Example 8H as a clear, viscous residue. MS: (CI, NH₃) (M+H)⁺ = 419.2 (100%).

5

Table 1d



Ex. No.	R ²²	R ²³	X	Y	stereo 2:3:4:5	Ki	IC ₉₀	MS (M+H)
7A			OH	OAc	RSSR	+	+++	449.1
7B					RSSR	-	++	465.2
7C					RSSR	+++	+++	453.2
7D					RSSR	+++	++	469.2
7E					RSSR	++	+++	449.2
7F					RSSR	+++	+++	481.2
7G					RSSR	++	+++	461
7H					RSSR	+	++	447.2

7I			RSSR	++	+++	477.2
7J		OAc OAc	RSSR	++	+	691.3
7K		OH OAc	RSSR	++	+	649.3
7L			RSSR	++	+	633.2
7M			RSSR	+++	+++	653.2
7N			RSSR	+	+	649.2
7O		OMs OH	RSSR	++	+++	685
7P			RSSR	+++	+++	661
7Q			RSSR	++	++	619
7R			RSSR	+	+	647.3
7S			RSSR		+	691
7T		OH H	RSSR	+++	+	591

7U



OH

H

RSSR

+++

+++

419

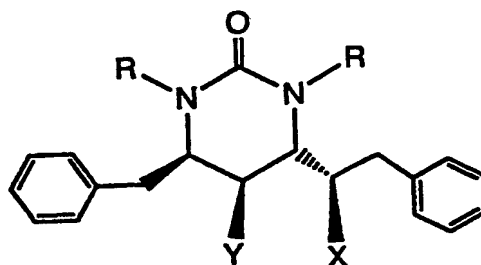
Table 1d (continued)

Ex. No.	R ²² R ²³	X	Y	<u>stereo</u> <u>2:3:4:5</u>	<u>HPLC</u> <u>K_i</u>	<u>MS</u> <u>(M+H)</u>
7AA		MEM	O=	RS-R	+	521
7AB		hydroxy	HO-N=	RS-R	+	448
7AC		hydroxy	MeO-N=	RS-R	++	462
7AD		oxazole	oxazole	R--R	++	442
7AE		methoxy	hydroxy	RSSR	++	449
7AF		methoxy	methoxy	RSSR	+	463
7AG		benzyloxy	benzyloxy	RSSR	+	615
7AH		benzyloxy	hydroxy	RSSR	+	525
7AI		allyloxy	allyloxy	RSSR		--
7AJ		allyloxy	hydroxy	RSSR	++	475


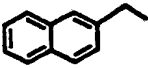
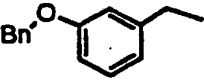
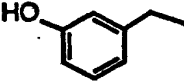
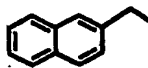

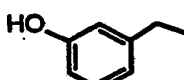
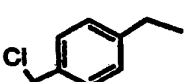
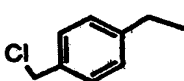
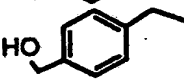
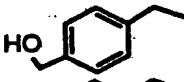
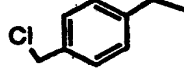
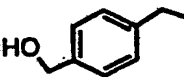
7AK 2-naphthyl OCH₂OH H RR-R +++ 621
 methyl

Table 1e

5



Ex. No.	R	X	Y	Ki	IC ₉₀	MS (M+H)
7V		F	OH	++	+++	437.26
7W		F	OH	++	++	409.23
7X		F	OH	++	+	609.29
7Y		N ₃	OH	++	+	632
7Z		OH	OH	++	+++	435.2
8A		OH	OMEM			523.4
8B		F	OH	++	+++	437.3

8C		F	OMEM	+	++	525.4
8D		F	OMEM	++		697.5
8E		F	OH	++		721
8F		F	OH	++ (HPLC)	+++	541.0
8G		H	OH	++ (HPLC)	++	591.5
8H		H	OH	+++ (HPLC)	+++	419.1
8I		H	OH	+++ (HPLC)	+++	523.2
8J		H	OH	++ (HPLC)		587.2
8K	 	H	OH	++ (HPLC)		569.2
8L	 	H	OH	++ (HPLC)		569.2
8M		H	OH	+++ (HPLC)		551

8N



F

H

+

+++

421.2

(HPLC)

5 Using the above-described techniques or variations thereon appreciated by those of skill in the art of chemical synthesis, the compounds of Tables 3-13 (shown below) can also be prepared.

10

Utility

 The compounds of formula (I) possess retroviral protease inhibitory activity and are therefore useful as antiviral agents for the treatment of viral diseases.

15 More particularly, the compounds of formula (I) possess HIV protease inhibitory activity and are effective as inhibitors of HIV growth. The protease inhibitory activity of the compounds of the present invention is demonstrated using standard assays of protease activity,

20 for example, using the assay described below for assaying inhibitors of HIV protease activity. The ability of the compounds of the present invention to inhibit viral growth or infectivity is demonstrated in standard assay of viral growth or infectivity, for

25 example, using the assays described below.

 A compound is considered to be active if it has an IC₅₀ or K_i value of less than about 1 mM for the inhibition of HIV protease or HIV viral growth or infectivity.

30

HIV Protease Inhibition Assay- Spectroscopic Method

Materials:

Protease: Inclusion bodies of E. coli harboring plasmid containing HIV protease under the control of an inducible T7 promoter were prepared according to Cheng et. al (1990) Gene 87: 243. Inclusion bodies were solubilized in 8 M urea, 50 mM tris pH 8.0. Protease activity was recovered by dilution 20-fold into buffer containing 50 mM sodium acetate pH 5.5, 1 mM EDTA, 10% glycerol and 5% ethylene glycol. Enzyme was used at a final concentration of 1.0-10 µg/ml.

Substrate: Peptide of sequence: Ala-Thr-His-Gln-Val-Tyr-Phe(NO₂)-Val-Arg-Lys-Ala, containing p-nitrophenylalanine (Phe(NO₂)), was prepared by solid phase peptide synthesis as previously described by Cheng et al. (1990) Proc. Natl. Acad. Sci. USA 87: 9660. Stock solutions of 10 mM were prepared in DMSO.

Inhibitory compounds were dissolved in sufficient DMSO to make 2.5 or 25 mM stock solutions. All further dilutions were done in DMSO.

Reactions:

Compound (1-5 µL) and HIV protease were mixed in buffer containing 50 mM MES, pH 6.5, 1 M NaCl, 1 mM EDTA, 1 mM dithiothreitol, 10% glycerol. Reactions were initiated by the addition of peptide substrate to a final concentration of 240 µM, and absorbance at 300 nm monitored for 10 min. Values of K_i for inhibitor binding were determined from percent activity measurements in the presence and absence of known concentration of inhibitor, using a value of 0.07 mM for the K_m of the substrate (Cheng et al. (1990) Proc. Natl. Acad. Sci. USA 87: 9660).

The HIV-1 protease inhibitory activity of representative compounds of the invention is shown in Table 1 and 2.

HIV Protease Inhibition Assay- HPLC Method5 Materials:

Protease: Inclusion bodies of E. coli harboring plasmid containing plasmid T1718R with a synthetic gene coding for a single-chain tethered dimer of HIV protease were prepared as described in Cheng et al. (Proc. Natl. Acad. Sci. USA, 87, 9660-9664, 1990). Active protease was prepared as described therein by extraction with 67% acetic acid, dilution 33-fold with water, dialysis against water and then against a "refolding buffer" consisting of 20 mM MES, 1 mM dithiothreitol and 10% glycerol. Protease was stored as a stock preparation at 10 μ M in refolding buffer.

Substrate: Peptide of sequence: aminobenzoyl-Ala-Thr-His-Gln-Val-Tyr-Phe(NO₂)-Val-Arg-Lys-Ala containing p-nitrophenylalanine, was prepared by solid phase synthesis as previously described Cheng et al., op.cit. Stock solutions of 2 mM substrate were prepared in DMSO.

Inhibitory compounds were dissolved in sufficient DMSO to make 3 mM stock solutions. All further dilutions were prepared in "assay buffer": 1 M NaCl, 50 mM MES, pH 5.5, 1 mM EDTA, 1mM DTT, 20% glycerol.

Reactions:

Enzyme reaction: In a 2 ml screw-cap centrifuge tube were added 50ul protease (final concentration 0.25 nM) and 0.1 ml inhibitory compound (final concentration 0.1-12,500). After 15 min preincubation at room temperature, the reaction was started with the addition of .05 ml substrate (final concentration 5 μ M). Incubation was carried out at 30 C. for 1 hr. The reaction was stopped with 1 ml 0.1 M ammonium hydroxide.

HPLC measurement of product formation: The product (aminobenzoyl-Ala-Thr-His-Gln-Val-Tyr) was separated from substrate on a Pharmacia MonoQ anion exchange column. The injection volume was 0.2 ml. The mobile phases were: A (20 mM trisHCl, pH 9.0, 0.02% sodium Azide, 10% acetonitrile), B (20 mM tris HCl, pH 9.0, 0.02% sodium azide, 0.5 M ammonium formate, 10% acetonitrile). The mobile phases were pumped at 1 ml/min, with a gradient from 0 to 30% B in 5 min, 100 % B for 4 min to wash the column, and a re-equilibration for 4 min. The retention time of the product was 3.6 min. Detection with a Shimadzu model RF535 fluorescence monitor was at 330 nm (excitation) and 430 (emission). The K_i was calculated from the formula $K_i = I / ((K_m + S - FA \cdot S) / (FA \cdot K_m)) - 1$ where I = inhibitory concentration, S = substrate concentration, FA = fractional activity = cm peak height with inhibitor/cm peak height without inhibitor, and K_m = Michaelis constant = 20 μ M.

20 HIV Yield Reduction Cell Assay

Materials: MT-2, a human T-cell line, was cultured in RPMI medium supplemented with 5% (v/v) heat inactivated fetal calf serum (FCS), L-glutamine and gentamycin. Human immunodeficiency virus strains, HIV (3B) and HIV(RF) were propagated in H-9 cells in RPMI with 5% FCS. Poly-L-lysine (Sigma) coated cell culture plates were prepared according to the method of Harada et al. (1985) Science 229: 563-566. MTT, 3-(4,5-dimethyl-thiazol-2yl)-2,5-diphenyltetrazolium bromide, was obtained from Sigma.

Method: Test compounds were dissolved in dimethylsulfoxide to 5 mg/mL and serially diluted into RPMI medium to ten times the desired final concentration. MT-2 cells (5×10^5 /mL) in 2.3 mL were mixed with 0.3 ml of the appropriate test compound

- solution and allowed to sit for 30 minutes at room temperature. HIV (3B) or HIV (RF) ($\sim 5 \times 10^5$ plaque forming units/mL) in 0.375 ml was added to the cell and compound mixtures and incubated for one hour at 36°C.
- 5 The mixtures were centrifuged at 1000 rpm for 10 minutes and the supernatants containing unattached virus were discarded. The cell pellets were suspended in fresh RPMI containing the appropriate concentrations of test compound and placed in a 36°C, 4% CO₂ incubator. Virus
- 10 was allowed to replicate for 3 days. Cultures were centrifuged for 10 minutes at 1000 rpm and the supernatants containing cell free progeny virus were removed for plaque assay.
- The virus titers of the progeny virus produced in
- 15 the presence or absence of test compounds were determined by plaque assay. Progeny virus suspensions were serially diluted in RPMI and 1.0 mL of each dilution was added to 9 ml of MT-2 cells in RPMI. Cells
- 20 and virus were incubated for 3 hours at 36°C to allow for efficient attachment of the virus to cells. Each virus and cell mixture was aliquoted equally to two wells of a six well poly-L-lysine coated culture plate and incubated overnight at 36°C, 4% CO₂. Liquid and
- 25 unattached cells were removed prior to the addition of 1.5 mL of RPMI with 0.75% (w/v) Seaplaque agarose (FMC Corp.) and 5% FCS. Plates were incubated for 3 days and a second RPMI/agarose overlay was added. After an additional 3 days at 36°C, 4% CO₂, a final overlay of
- 30 phosphate-buffered saline with 0.75% Seaplaque agarose and 1 mg MTT/mL was added. The plates were incubated overnight at 36°C. Clear plaques on a purple background were counted and the number of plaque forming units of virus was calculated for each sample. The antiviral
- 35 activity of test compounds was determined by the percent reduction in the virus titer with respect to virus grown in the absence of any inhibitors.

HIV Low Multiplicity Assay

Materials: MT-2, a human T-cell line, was cultured
5 in RPMI medium supplemented with 5% (v/v) heat
inactivated fetal calf serum (FCS), L-glutamine and
gentamycin (GIBCO). Human immunodeficiency virus
strains HIV (3B) and HIV (RF) were propagated in H-9
cells in RPMI with 5% FCS. XTT, benzene-sulfonic acid,
10 3,3'-[1-[(phenyl-amino)carbonyl]-3,4-tetrazolium]bis(4-
methoxy-6-nitro)-, sodium salt, was obtained from Starks
Associates, Inc.

Method: Test compounds were dissolved in dimethyl-
sulfoxide to 5 mg/ml and serially diluted into RPMI
15 medium to ten times the desired final concentration.
MT-2 cells (5×10^4 /0.1 mL) were added to each well of a
96 well culture plate and 0.02 mL of the appropriate
test compound solution was added to the cells such that
each compound concentration was present in two wells.
20 The cells and compounds were allowed to sit for 30
minutes at room temperature. HIV(3B) or HIV(RF) ($\sim 5 \times$
 10^5 plaque forming units/mL) was diluted in medium and
added to the cell and compound mixtures to give a
multiplicity of infection of 0.01 plaque forming
25 unit/cell. The mixtures were incubated for 7 days at
36°C, during which time the virus replicated and caused
the death of unprotected cells. The percentage of cells
protected from virus induced cell death was determined
by the degree of metabolism of the tetrazolium dye, XTT.
30 In living cells, XTT was metabolized to a colored
formazan product which was quantitated
spectrophotometrically at 450 nm. The amount of colored
formazan was proportional to the number of cells
protected from virus by the test compound. The
35 concentration of compound protecting either 50% (IC₅₀)

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or 90% (IC₉₀) with respect to an uninfected cell culture was determined.

The HIV inhibitory activity of representative compounds of the present invention in the whole cell infectivity assay described above is shown in Table 2.

Table 2

Example Number	IC ₉₀
1C	+++
1X	+++
1Z	+++
1AC	+++
1AF	+++
1AH	+++
1AQ	+++

The IC₉₀ values in Table 2 are indicated as: +++ = <10 µg/mL.

In the Tables herein the K_i values were determined using the assay conditions described above under HIV Protease Inhibition Assay. The K_i values are indicated as follows: +++ = <10 nM; ++ = 10 nM to 1 µM; + = >1 µM.

In the Tables herein the IC₉₀ values were determined using the assay conditions described above under HIV Low Multiplicity Assay. The IC₉₀ values are indicated as follows: +++ = <10 µg/mL; ++ = 10 to 100 µg/mL; + = >100 µg/mL.

Dosage and Formulation

The antiviral compounds of this invention can be administered as treatment for viral infections by any

means that produces contact of the active agent with the agent's site of action, the viral protease, in the body of a mammal. They can be administered by any conventional means available for use in conjunction with pharmaceuticals; either as individual therapeutic agents or in a combination of therapeutic agents. They can be administered alone, but generally administered with a pharmaceutical carrier selected on the basis of the chosen route of administration and standard pharmaceutical practice.

The dosage administered will, of course, vary depending upon known factors, such as the pharmacodynamic characteristics of the particular agent and its mode and route of administration; the age, health and weight of the recipient; the nature and extent of the symptoms; the kind of concurrent treatment; the frequency of treatment; and the effect desired. A daily dosage of active ingredient can be expected to be about 0.001 to 1000 milligrams per kilogram of body weight, with the preferred dose being 0.1 to about 30 mg/kg.

Dosage forms (compositions suitable for administration contain from about 1 milligram to about 100 milligrams of active ingredient per unit. In these pharmaceutical compositions the active ingredient will ordinarily be present in an amount of about 0.5-95% by weight based on the total weight of the composition.

The active ingredient can be administered orally in solid dosage forms, such as capsules, tablets, and powders, or in liquid dosage forms, such as elixirs, syrups, and suspensions. It can also be administered parenterally, in sterile liquid dosage forms.

Gelatin capsules contain the active ingredient and powdered carriers, such as lactose, starch, cellulose derivatives, magnesium stearate, stearic acid, and the like. Similar diluents can be used to make compressed

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tablets. Both tablets and capsules can be manufactured as sustained release products to provide for continuous release of medication over a period of hours.

- 5 Compressed tablets can be sugar coated or film coated to mask any unpleasant taste and protect the tablet from the atmosphere, or enteric coated for selective disintegration in the gastrointestinal tract.

- 10 Liquid dosage forms for oral administration can contain coloring and flavoring to increase patient acceptance.

- In general, water, a suitable oil, saline, aqueous dextrose (glucose), and related sugar solutions and glycols such as propylene glycol or polyethylene glycols are suitable carriers for parenteral solutions.
- 15 Solutions for parenteral administration preferably contain a water soluble salt of the active ingredient, suitable stabilizing agents, and if necessary, buffer substances. Antioxidizing agents such as sodium bisulfite, sodium sulfite, or ascorbic acid, either
- 20 alone or combined, are suitable stabilizing agents. Also used are citric acid and its salts and sodium EDTA. In addition, parenteral solutions can contain preservatives, such as benzalkonium chloride, methyl- or propyl-paraben, and chlorobutanol.

- 25 Suitable pharmaceutical carriers are described in Remington's Pharmaceutical Sciences, Mack Publishing Company, a standard reference text in this field.

- Useful pharmaceutical dosage-forms for administration of the compounds of this invention can be
- 30 illustrated as follows:

Capsules

- A large number of unit capsules are prepared by filling standard two-piece hard gelatin capsules each
- 35 with 100 milligrams of powdered active ingredient, 150

milligrams of lactose, 50 milligrams of cellulose, and 6 milligrams magnesium stearate.

Soft Gelatin Capsules

5 A mixture of active ingredient in a digestable oil such as soybean oil, cottonseed oil or olive oil is prepared and injected by means of a positive displacement pump into gelatin to form soft gelatin capsules containing 100 milligrams of the active
10 ingredient. The capsules are washed and dried.

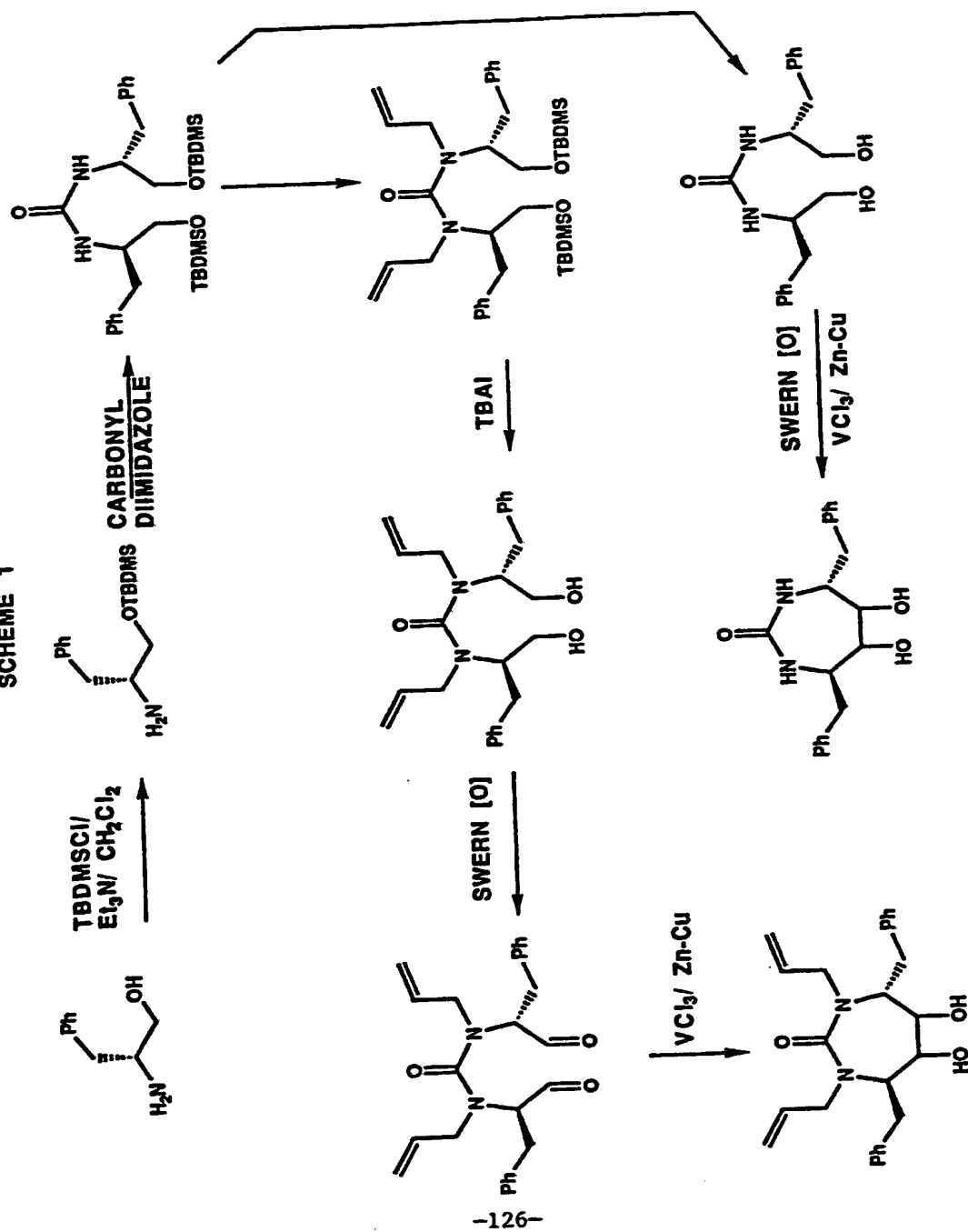
Tablets

 A large number of tablets are prepared by conventional procedures so that the dosage unit was 100
15 milligrams of active ingredient, 0.2 milligrams of colloidal silicon dioxide, 5 milligrams of magnesium stearate, 275 milligrams of microcrystalline cellulose, 11 milligrams of starch and 98.8 milligrams of lactose. Appropriate coatings may be applied to increase
20 palatability or delay absorption.

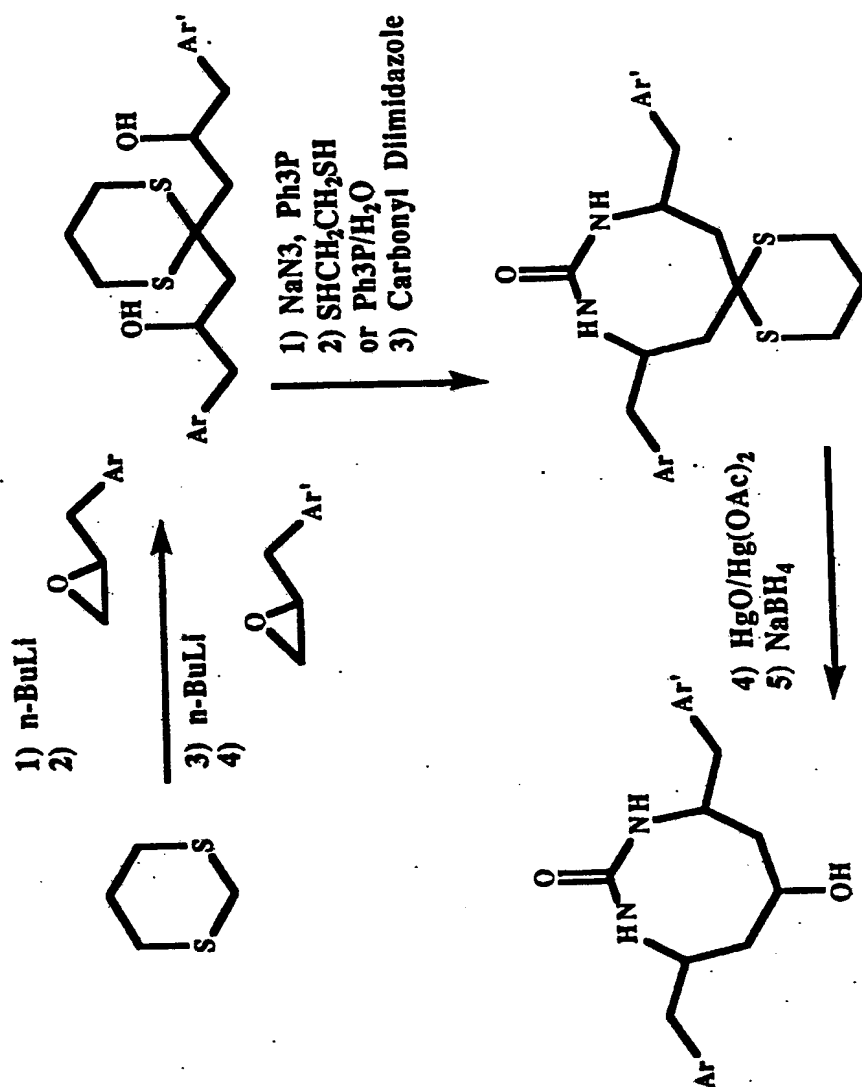
 Tables 3-16 below set forth representative compounds of the present invention.

25 In Tables 3-12, each of the Examples 1-5763 includes all stereoisomers of the indicated structure.

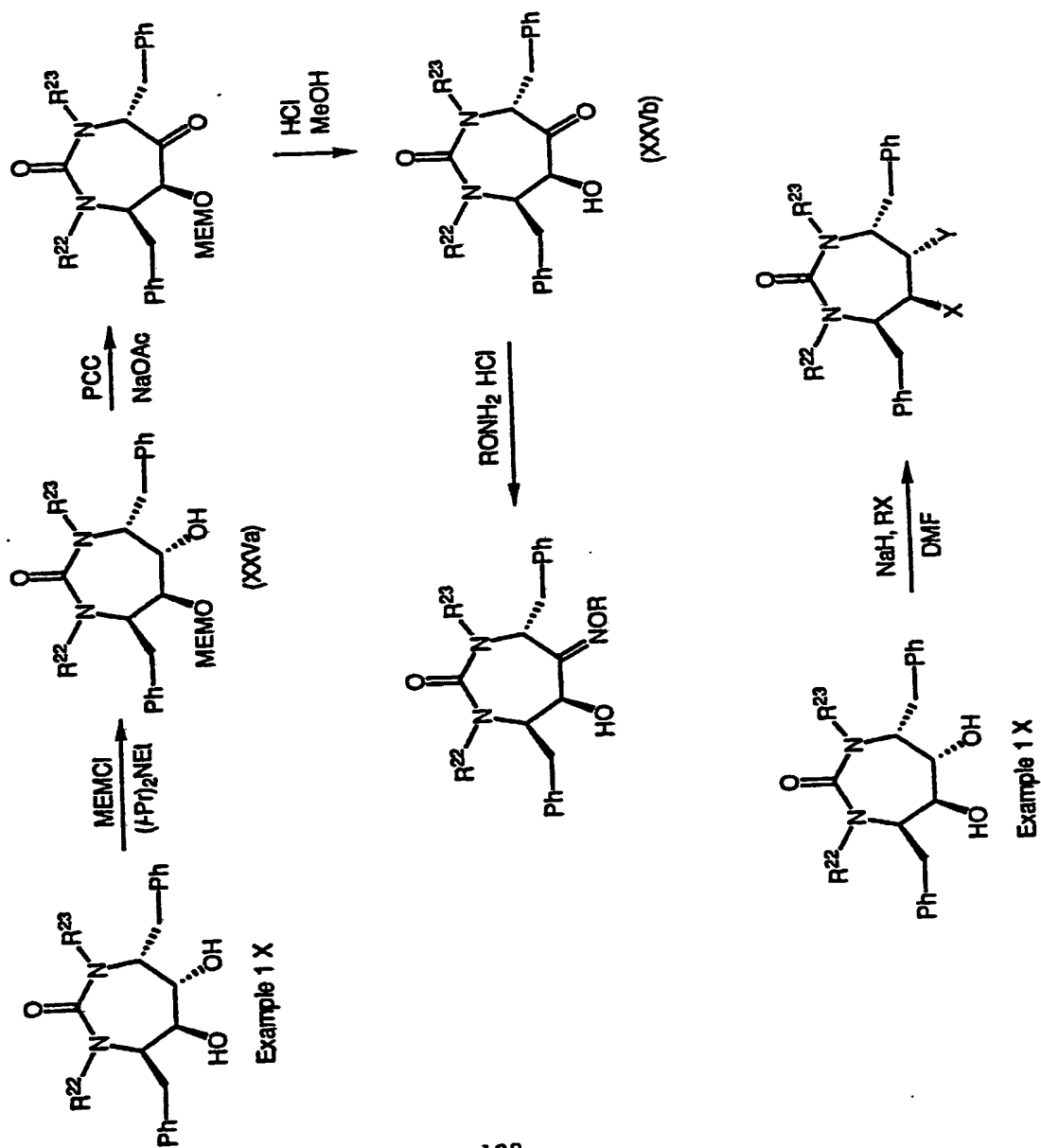
SCHEME 1



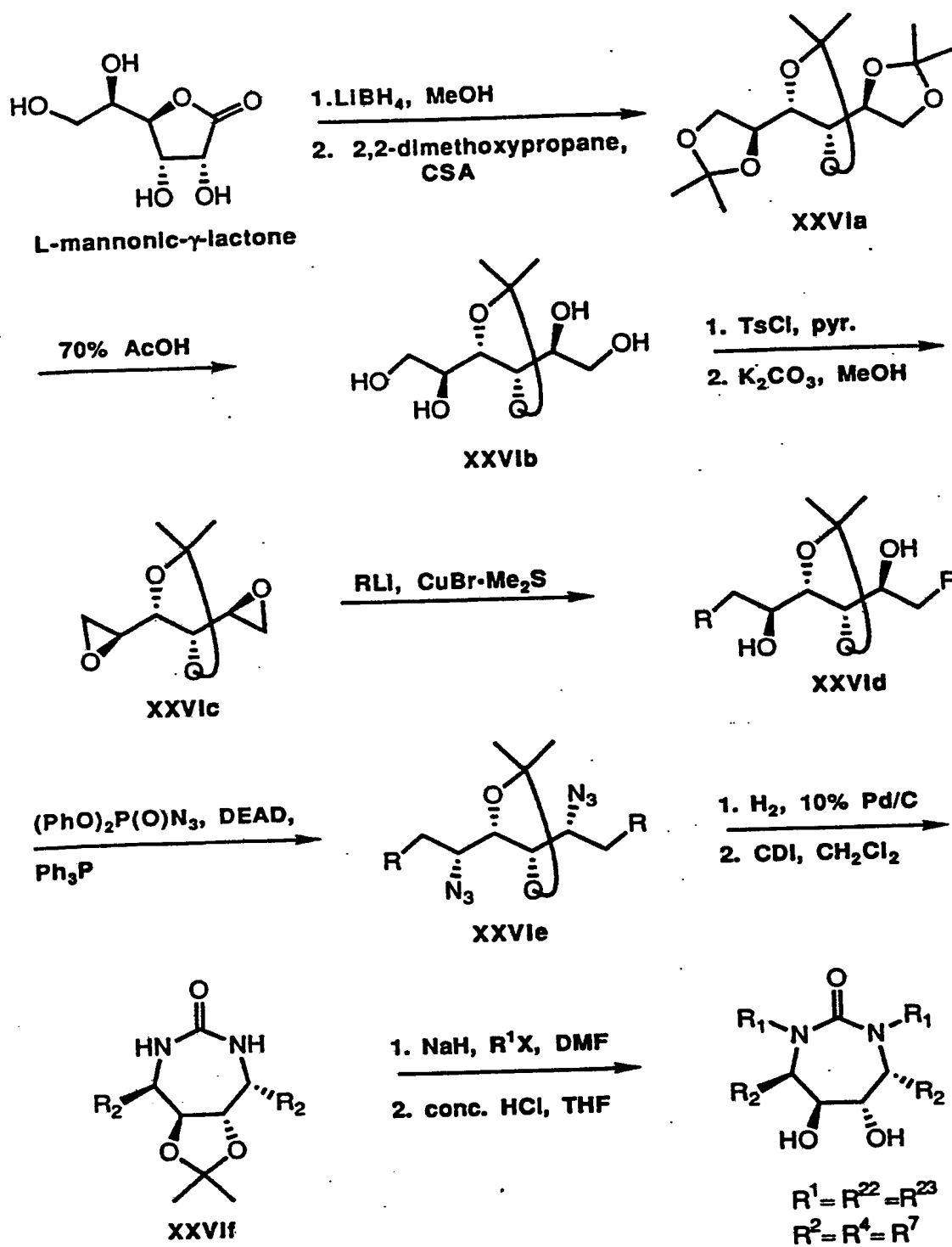
Scheme 2



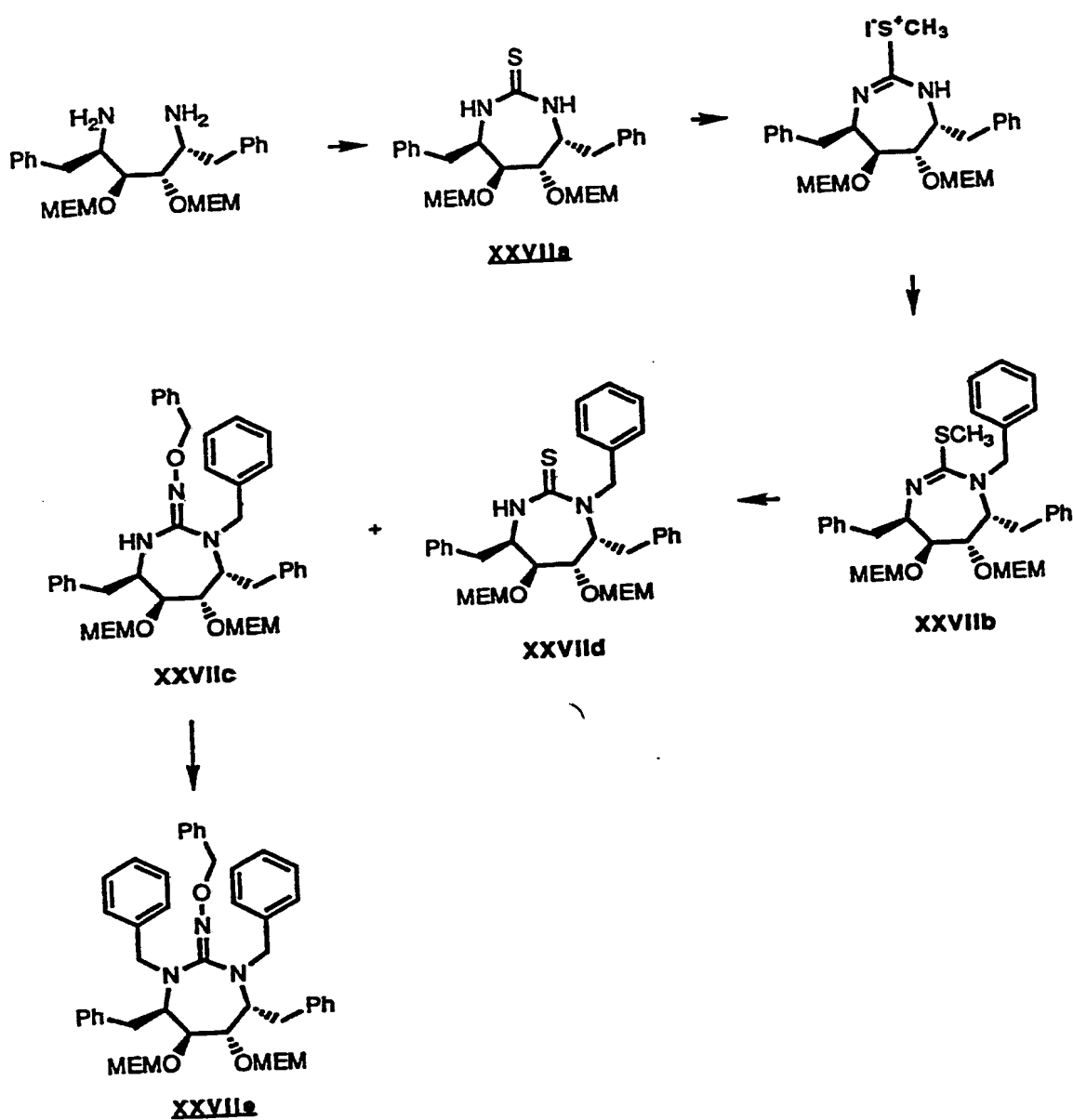
Scheme 3



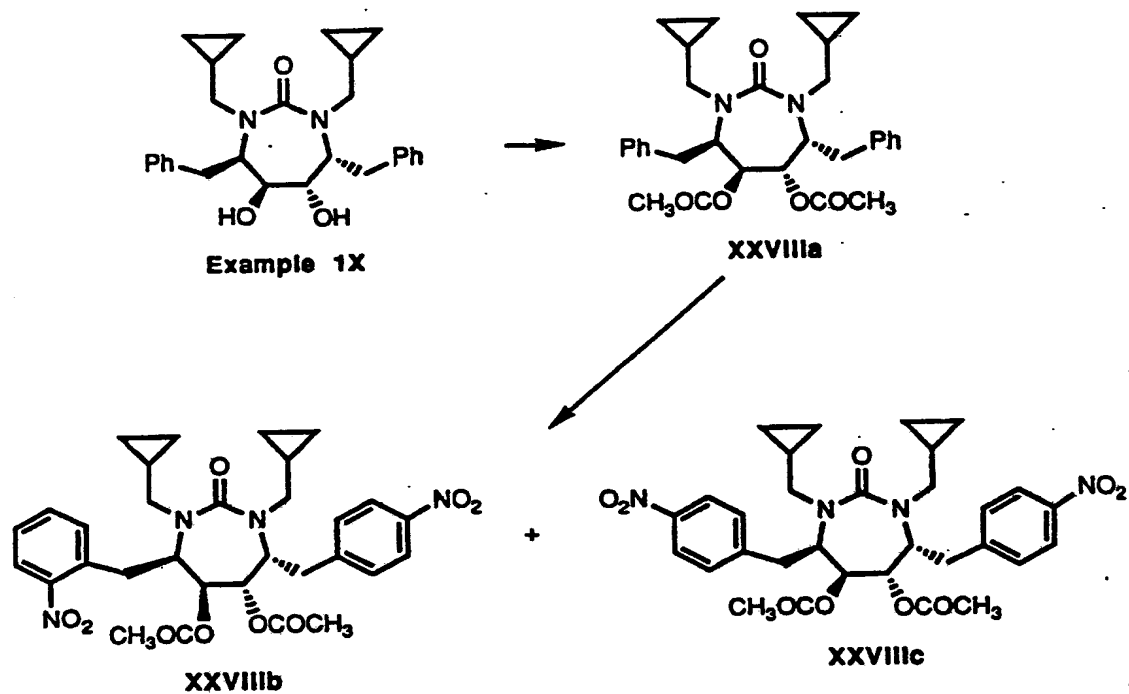
Scheme 4



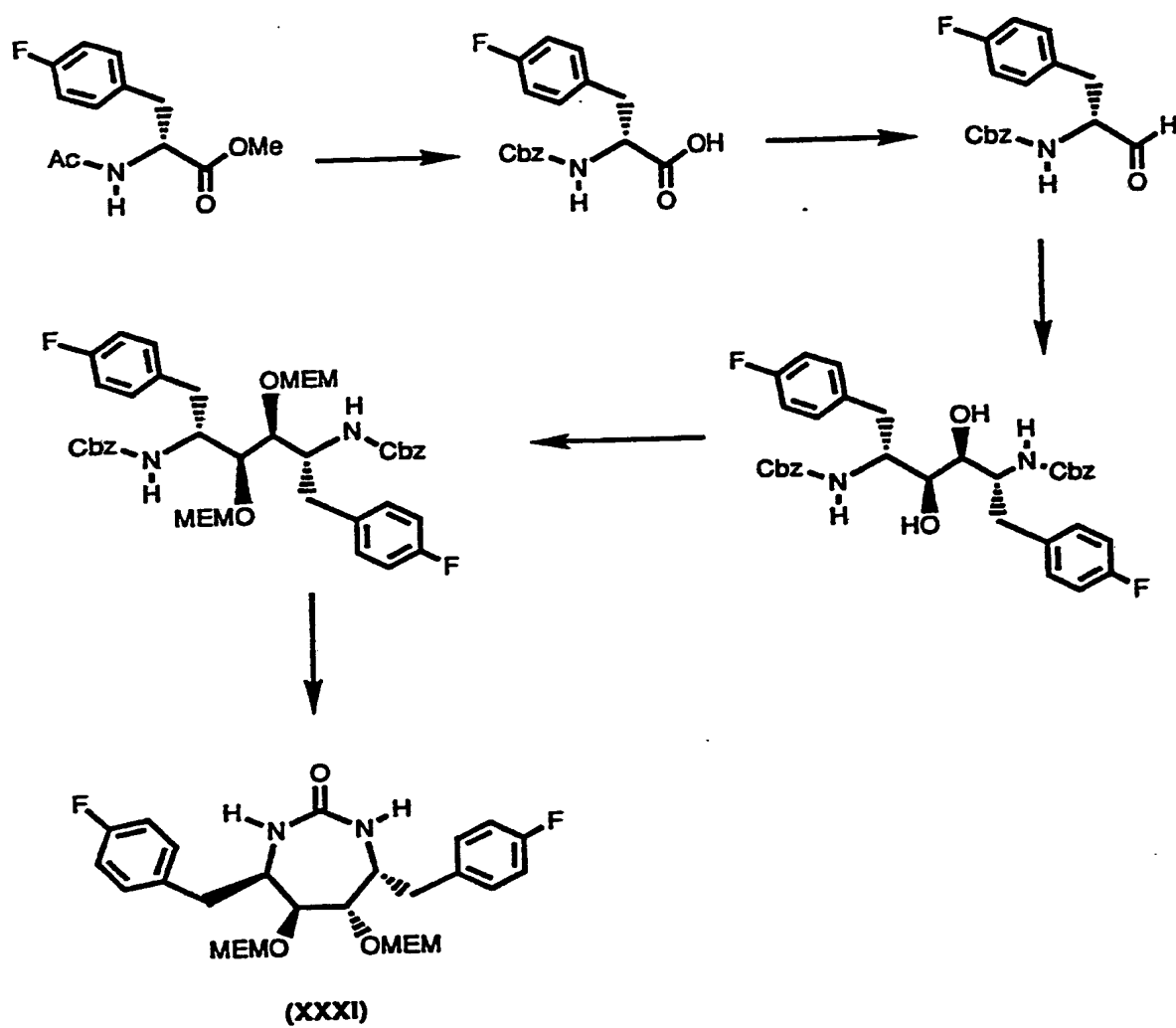
Scheme 5



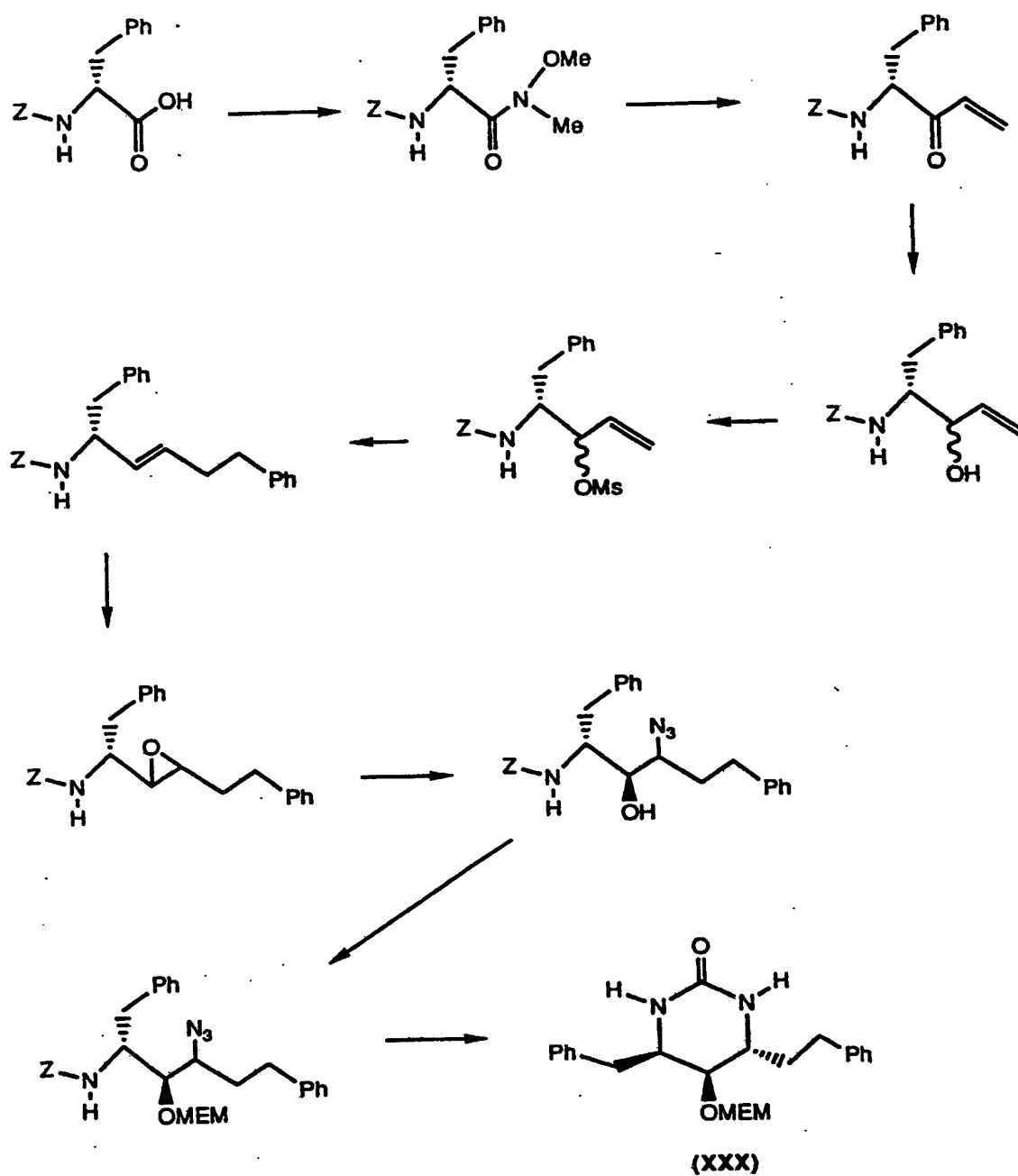
Scheme 6



Scheme 7



Scheme 8



Scheme 9

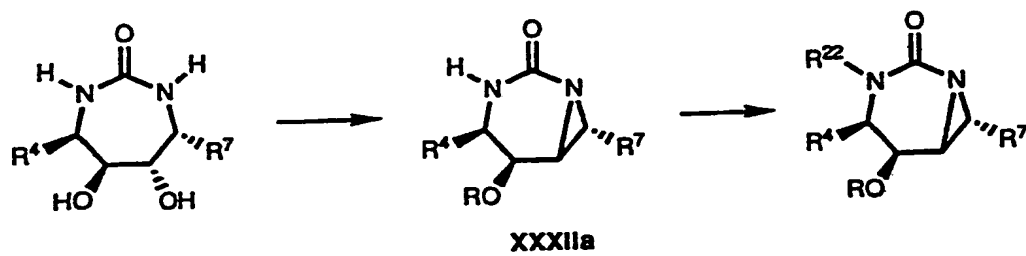
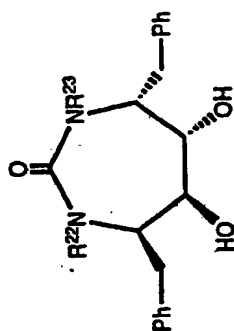


Table 2A




Example Number	R22	R23	MS (M+H)	Ki	IC90	m.p. (°C)
2A	CH ₂ CHC(CH ₃) ₂	CH ₂ CHC(CH ₃) ₂	463	+++	+++	165.1
2B	CH ₂ CH ₂ CH ₂ C ₆ H ₅	CH ₂ CH ₂ CH ₂ C ₆ H ₅	563	++	+++	120.7
2C	CH ₂ CHCH ₂	CH ₂ CHC(CH ₃) ₂	435	+++	+++	62
2D	CH ₂ CHC(CH ₃) ₂	CH ₃	409	++		sin 65 108
2E	CH ₂ C ₆ H ₄ P-F	CH ₂ C ₆ H ₄ P-F	543	+++	+++	
2F		CH ₂ C ₆ H ₅	471	+++	+++	
2G	CH ₂ CHCH ₂	CH ₂ C ₆ H ₅	457	+++	+++	226.5
2H	CH ₂ C ₆ H ₅	CH ₃	431	++	+++	81

Table 2A (continued).


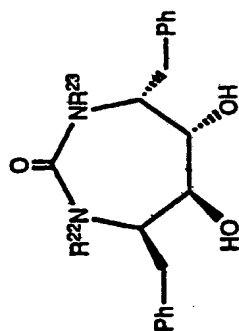
2I	CH ₂ CHC(CH ₃) ₂		440	+++	+++	76
2J	CH ₂ C ₆ H ₄ m-OCF ₃	CH ₂ C ₆ H ₄ m-OCF ₃	675	++	+++	128.3
2K	CH ₂ C ₆ H ₃ m-CF ₃ p-F	CH ₂ C ₆ H ₃ m-CF ₃ p-F	679	++	+++	167.2
2L	CH ₂ C ₆ H ₄ pOCF ₃	CH ₂ C ₆ H ₄ pOCF ₃	675	++	+++	136.8
2M	CH ₂ C ₆ H ₃ m-Fm-CF ₃	CH ₂ C ₆ H ₃ m-Fm-CF ₃	679	++		162.3

Table 2B



Example Number	R ²²	R ²³	MS (M+H)	Ki	IC ₉₀	Notes
2N	C ₆ H ₅ CH=CHCH ₂	C ₆ H ₅ CH=CHCH ₂	559.30	+++	+++	1
2O	p-CH ₃ O-C ₆ H ₄ CH ₂	p-CH ₃ O-C ₆ H ₄ CH ₂	567.29	++	+++	1
2P	(CH ₃) ₂ NCH ₂ CH ₂ -	(CH ₃) ₂ NCH ₂ CH ₂	468.62	+	+	1
2Q	C ₆ H ₅ CH ₂	H	417.22	+++	+++	1
2R	m-F-C ₆ H ₄ CH ₂	m-F-C ₆ H ₄ CH ₂	543.25	+++	+++	1
2S	o-F-C ₆ H ₄ CH ₂	o-F-C ₆ H ₄ CH ₂	543.25	++	+++	1
2T	m-CH ₃ O-C ₆ H ₄ CH ₂	m-CH ₃ O-C ₆ H ₄ CH ₂	567.29	+++	+++	1
2U	3-F, 4-F-C ₆ H ₄ CH ₂	3-F, 4-F-C ₆ H ₄ CH ₂	579.23	++	+++	1
2V	p-CH ₃ -C ₆ H ₄ CH ₂	p-CH ₃ -C ₆ H ₄ CH ₂	535.30	+++	+++	1

Table 2B (continued)



2W	p-Cl-C ₆ H ₄ CH ₂	p-Cl-C ₆ H ₄ CH ₂	575.19	+++	1
2X		pF-C ₆ H ₄ CH ₂	489.26	+++	2
2Y	p-CF ₃ -C ₆ H ₄ CH ₂	p-CF ₃ -C ₆ H ₄ CH ₂	643.24	+++	1
2Z	m-Cl-C ₆ H ₄ CH ₂	m-Cl-C ₆ H ₄ CH ₂	575.19	+++	1
3A	m-CF ₃ -C ₆ H ₄ CH ₂	m-CF ₃ -C ₆ H ₄ CH ₂	643.24	+	1
3B	m-NO ₂ -C ₆ H ₄ CH ₂	m-NO ₂ -C ₆ H ₄ CH ₂	597.23	+++	1
3C	m-CH ₃ -C ₆ -H ₄ CH ₂	m-CH ₃ -C ₆ -H ₄ CH ₂	535.30	+++	1
3D	o-CH ₃ -O-C ₆ H ₄ CH ₂	o-CH ₃ -O-C ₆ H ₄ CH ₂	567.29	++	1
3E	o-CH ₃ -O-C ₆ H ₄ CH ₂	o-CH ₃ -O-C ₆ H ₄ CH ₂	579.23	+++	1
3F	o-Cl-C ₆ H ₄ CH ₂	o-Cl-C ₆ H ₄ CH ₂	575.58	+++	1
3G	m-Br-C ₆ H ₄ CH ₂	m-Br-C ₆ H ₄ CH ₂	665.00	+++	1
3H	p-F-C ₆ H ₄ CH ₂	H	435.21	+++	1
3I	p-Br-C ₆ H ₄ CH ₂	p-Br-C ₆ H ₄ CH ₂	663.09	++	1

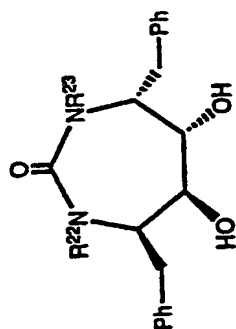
Table 2B (continued)

3J		m-Cl-C ₆ H ₄ CH ₂	505.23	+++	2
3K		m-NH ₂ -C ₆ H ₄ CH ₂	537.28	+++	3,4

NOTES:

1. Prepared according to the general alkylation procedure (Procedure 5). Yields of monoalkyl compounds were favored by using one equivalent of alkylating agent.
2. Prepared by alkylating the appropriate monoalkyl compound.
3. Isolated as the dihydrochloride salt.
4. Preparation: a solution of 41.3 mg of Ex.3B was dissolved in a mixture of 10ml of ethanol and 1 ml of 1 N hydrochloric acid. Catalyst (20 mg of 10% palladium on carbon) was added, and the mixture was hydrogenated at atm pressure for 16 hours. A quantitative yield of Ex. 3K as isolated as the hydrochloride salt.

Table 2C



Example Number	Stereo-isomer	R ²²	R ²³	K1	IC ₅₀	MP (°C)	(M+1, 100%)
3L	RSSR	CH ₂ C≡CH	CH ₂ C≡CH	++		196-197	507
3M	RSSR	2-picolinyl	2-picolinyl	++		151-153	509
3N	RSSR	CH ₂ CH ₂ OCH ₃	CH ₂ CH ₂ OCH ₃	++		183-185	443
3O	RSSR	CH ₂ C ₆ H ₁₂	CH ₂ C ₆ H ₁₂	++		-	519
3P	RSSR	CH ₂ CH ₂ C(CH ₃) ₃	CH ₂ CH ₂ C(CH ₃) ₃	++		242-245	495
3Q	RSSR	CH ₂ C ₆ H ₆	CH ₂ C ₆ H ₆	++		-	417
3R	RSSR	CH ₂ CH ₂ OCH ₂ CH ₃	CH ₂ CH ₂ OCH ₂ CH ₃	+		148-150	471
3S	RSSR	3-methyl-5-oxazoliny1-methyl	H	++		-	464
3T	RSSR	1-naphthylmethyl	1-naphthylmethyl	++		231-233	607 (85%)

Table 2C (continued)

3U	RSSR	2-naphthylmethyl	2-naphthylmethyl	+++	+++	202-204	607
3V	RSSR	n-butyl	benzyl	+++	+++	89-91	473
3W	RSSR	CH ₂ CH=CH ₂	cyclopropylmethyl	+++	+++	173-175	421
3X	RSSR	n-butyl	cyclopropylmethyl	+++	+++	-	437
3Y	RSSR	CH ₂ CH=C(CH ₃) ₃	benzyl	+++	+++	173-174	485
3Z	RSSR	3-methyloxymethyl	3-methyloxymethyl	+		-	527
4A	RSSR	benzyl	ethyl	+++	+++	190-193	445
4B	RSSR	benzyl	4-picolinyl	+++	+++	-	508
4C	RSSR	cyclopropylmethyl	4-picolinyl	+++	+++	-	472
4D	RSSR	CH ₂ CH ₂ OCH=CH ₂	CH ₂ CH ₂ OCH=CH ₂	++		-	467
4E	RSSR	benzyl	cyclopentylmethyl	+++	+++	198-200	499
4F	RSSR	cyclopropylmethyl	cyclopentylmethyl	+++	+++	-	463
4G	RSSR	benzyl	n-propyl	+++	+++	-	459
4H	RSSR	benzyl-(D ₇)	benzyl-(D ₇)	+++	+++	-	521

Table 2C (continued)

4I	RSSR	cyclopropylmethyl	cinnamyl	+++	+++	-	497
4J	RSSR	2,3,4,5,6-penta-fluoro-benzyl	2,3,4,5,6-penta-fluoro-benzyl	+	+++	193-195	687
4K	RSSR	cyclopropylmethyl	2-naphthylmethyl	+++	+++	175-178	521
4L	RSSR	cyclopentylmethyl	2-naphthylmethyl	+++	+++	89-92	549
4M	RSSR	benzyl	2-naphthylmethyl	+++	+++	90-92	557
4N	RSSR	cyclopropylmethyl	2-picolinyl	+++	+++	220-222	472
4O	RSSR	benzyl	2-quinolinylmethyl	++	+++	-	558
4P	RSSR	3-cyanobenzyl	3-cyanobenzyl	+++	+++	225-227	557
4Q	RSSR	3-benzoyloxybenzyl	3-benzoyloxybenzyl	++	+++	140-141	691
4R	RSSR	4-phenylbenzyl	4-phenylbenzyl	++	+++	109-110	659
4S	RSSR	3-allyl	2-naphthylmethyl	+++	+++	72-74	507
4T	RSSR	n-propyl	2-naphthylmethyl	+++	+++	150-152	509
4U	RSSR	n-butyl	2-naphthylmethyl	+++	+++	78-80	523
4V	RSSR	H	2-naphthylmethyl	+++	+++	210-213	467
4W	RSSR	2-adamantylethyl	2-adamantylethyl	++	+++	292-294	651

Table 2C (continued)

4X	RSSR	H	cyclopropylmethyl	++	189-190	381
4Y	RSSR	2-picolinyl	2-naphthylmethyl	++	+++ 165-166	558
4Z	RSSR	4-picolinyl	2-naphthylmethyl	+++	+++ 118-121	558
5A	RSSR	3-allyl	H	+	180-182	367
5B	RSSR	3-allyl	cyclopentylmethyl	+++	+++ 190-191	449
5C	RSSR	3-allyl	2-picolinyl	++	+++ 160-162	458
5D	RSSR	3-allyl	2-quinolinylmethyl	+++	+++ 145-146	507
5E	RSSR	3-allyl	4-picolinyl	++	+++	458
5F	RSSR	3-benzyloxybenzyl	3-benzyloxybenzyl	++	165-166	

Table 2C (continued)

5G	RSSR	3-picolinyl	cyclopropylmethyl	+++	+++	193-195	472
5H	RSSR	3-picolinyl	2-naphthylmethyl	+++	+++	94-96	558
5I	RSSR	3-hydroxybenzyl	3-hydroxybenzyl	+++	+++	101-103	539
5J	RSSR	vinylbenzyl	vinylbenzyl	+++	+++	158-160	559
5K	RSSR	3-cyclopropyl-methoxy-benzyl	3-cyclopropyl-methoxy-benzyl	+	+	170-172	697
5L	RSSR	3-allyloxybenzyl	3-allyloxybenzyl	+++	+++	170-172	619
5M	RSSR	3-allyloxybenzyl	3-hydroxybenzyl	+++	+++	78-79	579
5N	RSSR	3-ethoxybenzyl	3-ethoxybenzyl	++	++	185-186	595
5O	RSSR	3-picolinyl	3-picolinyl	+++	+++	-	509
5P	RSSR	4-benzyoxybenzyl	4-benzyoxybenzyl	+	+	80-81	719
5Q	RSSR	2-naphthylmethyl-	4-fluorobenzyl	+++	+++	105-107	575
5R	RSSR	4-hydroxybenzyl	4-hydroxybenzyl	+++	+++	115-117	539 (20%)
5S	RSSR	3-hydroxymethyl-benzyl	3-hydroxymethyl-benzyl	+++	+++	94-95	567 (40%)
5T	RSSR	3-carbomethoxy-benzyl	3-carbomethoxy-benzyl	+++	+++	-	623 (40%)

Table 2C (continued)

5U	RSSR	4-hydroxy- methylbenzyl	4-hydroxy- methylbenzyl	+++	+++	90-95	567
5V	RSSR	3-formylbenzyl	3-formylbenzyl	+++	+++	175-176	563 (30%)
5W	RSSR	4-cyanobenzyl	4-cyanobenzyl	+++	+++	140-143	557 (15%)
5X	RSSR	4-formylbenzyl	4-formylbenzyl	+++	+++	90-91	563 (52%)
5Y	RSSR	4-hydroxybenzyl	2-propyl	+++	+++	246-248	475 (80%)
5Z	RSSR	3-hydroxybenzyl	2-propyl	+++	+++	212-213	475 (90%)
6A	RSSR	3-carboxybenzyl	3-carboxybenzyl	+++	+++	190-195	189
6B	RSSR	4-carboxybenzyl	4-carboxybenzyl	++	++	210-211	189
6C	RSSR	3-formaldoxime- benzyl	3-formaldoxime- benzyl	+++	+++	176-178	593 (2%)

Table 2C (continued)

6D	RSSR	cyclopropylmethyl	3-hydroxybenzyl	+++	+++	233-234	486
6E	RSSR	cyclopropylmethyl	4-hydroxybenzyl	+++	+++	234-236	486
6F	RSSR	5-chloro-2-thienyl-methyl	5-chloro-2-thienyl-methyl	++	+++		587
6G	RSSR	cyclobutylmethyl	cyclobutylmethyl	+++	+++		463
6H	RSSR	cyclopentylmethyl	cyclopentylmethyl	+++	+++		491
6I	RSSR	n-butyl	CH ₂ CH-C(CH ₃) ₂	+++	+++		
6J	RSSR	n-butyl	cyclopentylmethyl	+++	+++		465
6K	RSSR	2-quinolinylmethyl	2-quinolinylmethyl	++			609
6L	RSSR	2-propyl	2-picolinyll	++	+++		460
6M	RSSR	p-CH ₃ OC ₆ H ₄ CH ₂ -	p-CH ₃ OC ₆ H ₄ CH ₂ -				567.29
6N	RSSR	(CH ₃) ₂ NCH ₂ CH ₂ -	(CH ₃) ₂ NCH ₂ CH ₂ -				469.32
6O	RSSR	C ₆ H ₅ CH ₂ -	H				417.22
6P	RSSR	o-F-C ₆ H ₄ CH ₂ -	o-F-C ₆ H ₄ CH ₂ -				543.25

Table 2C (continued).


6Q	RSSR	m-CH ₃ O-C ₆ H ₄ CH ₂ -	m-CH ₃ O-C ₆ H ₄ CH ₂ -	567.29
6R	RSSR	(CH ₃) ₂ NCH ₂ CH ₂ -	(CH ₃) ₂ NCH ₂ CH ₂ -	469.32
6S	RSSR	m,p-F ₂ -C ₆ H ₄ CH ₂ -	m,p-F ₂ -C ₆ H ₄ CH ₂ -	579.23
6T	RSSR	p-CH ₃ -C ₆ H ₄ CH ₂ -	p-CH ₃ -C ₆ H ₄ CH ₂ -	535.30
6U	RSSR	p-Cl-C ₆ H ₄ CH ₂ -	p-Cl-C ₆ H ₄ CH ₂ -	575.19 (35Cl)
6V	RSSR	p-F-C ₆ H ₄ CH ₂ -		183.4 489.26
6W	RSSR	p-CF ₃ -C ₆ H ₄ CH ₂ -	p-CF ₃ -C ₆ H ₄ CH ₂ -	643.24
6X	RSSR	m-Cl-C ₆ H ₄ CH ₂	m-Cl-C ₆ H ₄ CH ₂	210.3 575.19 (35Cl)
6Y	RSSR	m-CF ₃ -C ₆ H ₄ CH ₂	m-CF ₃ -C ₆ H ₄ CH ₂	643.24
6Z	RSSR	m-NO ₂ -C ₆ H ₄ CH ₂ -	m-NO ₂ -C ₆ H ₄ CH ₂ -	248.3 597.23
7A	RSSR	m-CH ₃ -C ₆ H ₄ CH ₂ -	m-CH ₃ -C ₆ H ₄ CH ₂ -	535.30
7B	RSSR	o-CH ₃ O-C ₆ H ₄ CH ₂ -	o-CH ₃ O-C ₆ H ₄ CH ₂ -	567.29
7C	RSSR	m,m-F ₂ -C ₆ H ₄ CH ₂ -	m,m-F ₂ -C ₆ H ₄ CH ₂ -	167.9 579.23

Table 2C (continued)


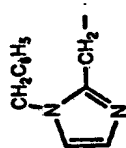
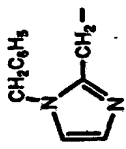
7D	RSSR	o-Cl-C ₆ H ₄ CH ₂ -	o-Cl-C ₆ H ₄ CH ₂ -	575.56 (³⁵ Cl)
7E	RSSR	m-Br-C ₆ H ₄ CH ₂ -	m-Br-C ₆ H ₄ CH ₂ -	210.7 665
7F	RSSR	p-F-C ₆ H ₄ CH ₂ -	H	435.21
7G	RSSR	p-Br-C ₆ H ₄ CH ₂ -	p-Br-C ₆ H ₄ CH ₂ -	663.09 (⁷⁹ Br)
7H	RSSR	m-Cl-C ₆ H ₄ CH ₂ -		126.0 505.23 (³⁵ Cl)
7I	RSSR	m-NH ₂ C ₆ H ₄ CH ₂ - (HCl)	m-NH ₂ C ₆ H ₄ CH ₂ - (HCl)	537.29
7J	RSSR	m,m'-Cl ₂ -C ₆ H ₄ CH ₂ -	m,m'-Cl ₂ -C ₆ H ₄ CH ₂ -	231.1 642.10 (³⁵ Cl)
7K	RSSR	m-(NH ₂ CH ₂)- C ₆ H ₄ CH ₂ -	m-(NH ₂ CH ₂)- C ₆ H ₄ CH ₂ -	565.32 ++
7L	RSSR	m-NO ₂ C ₆ H ₄ CH ₂ -	H	462.20
7M	RSSR	m-(NHCHO)- C ₆ H ₄ CH ₂ -	m-(NHCHO)-C ₆ H ₄ CH ₂ -	593.28

Table 2C (continued).

7N	RSSR	m-(NHCOCH ₃)- C ₆ H ₄ CH ₂ -	m-(NHCOCH ₃)- C ₆ H ₄ CH ₂ -	621.31
7o	RSSR	m, p-(HO) 2- C ₆ H ₄ CH ₂ -	m, p-(HO) 2-C ₆ H ₄ CH ₂ -	571.24
7p	RSSR	CH ₂ C ₆ H ₅ 	CH ₂ C ₆ H ₅ 	667.34

All of the compounds of Table 2C were synthesized using procedure 5.

Synthesis of compounds of the invention in Table 2d is described in further detail below.

Example 9A

5 To a solution of bis m-(HO-N=CH)-C₆H₄CH₂ cyclic urea (0.297, 0.502mmol) in methanol (5ml) was added borane-pyridine complex (0.464g, 5.02mmol) at -10°C and the resultant mixture was stirred for 15 minutes.

10 After treated with 4M HCl in dioxane (5ml), the reaction mixture was allowed to stir to room temperature for additional 1.5 hours. The solution was neutralized with sat. NaHCO₃ to pH=8, washed with water and dried over MgSO₄. After removal of solvents, the

15 residue was purified on silica gel plate with ethyl acetate: dichloromethane : methanol (50:50:2) to give 60mg of a solid, M.P. 214-216°C. ¹H NMR (CD₃OD): δ 8.03(s, 1H), 7.46 - 7.06 (m, 18H), 4.74 (d, J=13.9 Hz, 1H), 4.73 (d J=13.9Hz, 1H), 3.92(s, 2H), 3.62-3.59 (bs, 4H), 3.07-2.91(m, 6H) ¹³C NMR (CD₃OD): 163.88, 149.70,

20 141.26, 141.23, 139.92, 139.39, 138.73, 134.91, 131.50, 131.41, 130.64, 130.00 129.72, 129.64, 129.57, 129.53, 128.47, 127.45 127.42, 127.31, 71.96, 71.92, 67.42, 67.11, 58.65, 57.09, 57.03, 33.62, 33.57, MS: 594

25 (100%) 595 (M+H, 60%).

Example 9C

To a stirred solution of bis m-(HO-N=CH)-C₆H₄CH₂ cyclic urea (257 mg, 0.434 mmol), sodium cyano-

30 borohydride (290 mg, 4.6mmol) and trace amount of methyl orange in methanol (10ml) at room temperature was added dropwise 2N HCl at a rate sufficient to maintain a pH of 3-4 over 3 hours. The methanol was

35 removed by rotary evaporation. The residue was purified on a reverse phase TLC plate with 90% methanol

in water to give the product. ^1H NMR (CD_3OD): δ 7.33-7.09 (m, 18H), 4.75 (d, $J=13.9\text{Hz}$, 2H), 3.93 (s, 4H), 3.61- 3.56 (m, 4H), 3.06- 2.94 (m, 6H); ^{13}C NMR (CD_3OD): 162.42, 139.72, 137.88, 137.79, 129.92, 129.20, 128.31, 128.14, 128.06, 126.02, 70.36, 65.49, 57.40, 55.59, 32.09.

Example 9E

A solution of Example 9C (30mg) in methanol was treated with 4M HCl in dioxane at room temperature. All solvents were removed under vacuum to give hydroxylamine hydrochloride. ^1H NMR (CD_3OD): δ 7.44-7.23 (m, 4H), 6.99 (d, $J=6.2\text{Hz}$, 4H), 4.65 (d, $J=14.3\text{Hz}$, 2H), 4.35 (s, 4H), 3.70-3.66 (m, 4H), 3.12- 3.05 (m, 4H), 2.89-2.85 (m, 2H); ^{13}C NMR (CD_3OD): 163.68, 141.13, 140.39, 123.09, 132.03, 131.06, 130.59, 130.50, 129.59, 127.48, 71.82, 68.23, 57.40, 56.12, 33.70.

Example 9B

An unpurified sample of example 9C was purified on TLC plate using acetic acid: ethyl acetate: dichloromethane (5: 50:45) to give the acetic acid salt. ^1H NMR (CD_3OD): δ 7.35-7.08 (m, 18H), 4.73 (d, $J=10.6\text{Hz}$, 2H), 3.97 (s, 4H), 3.62-3.53 (m, 4H), 3.06-2.88 (m, 6H), 1.97 (s, 3H).

Example 9D

A solution of bis (m-CHO- $\text{C}_6\text{H}_4\text{CH}_2$) cyclic urea (119 mg, 0.212 mmol) and O-benzylhydroxylamine hydrochloride (203mg, 1.27mmol) in pyridine/ethanol (6ml 1:1) was refluxed for 3 hours. After removal of solvent, the residue was purified on T.L.C. plate with 15% ethyl acetate in dichloromethane to give the product (164mg)

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as a solid, M.P. 170.5-171°C. ^1H NMR(CDCl_3): δ 8.01 (s, 2H), 7.39- 7.07 (m, 28H), 5.09 (s, 4H), 4.81 (d, $J=14.3\text{Hz}$, 2H), 3.59 (bs, 2H), 3.52 (d, $J=11.0\text{Hz}$, 2H), 3.07- 2.88 (m, 8H); ^{13}C NMR(CDCl_3): 162.14, 148.55, 139.40, 138.62, 137.39, 132.50, 130.50, 129.43, 128.87, 128.62, 128.35, 128.27, 128.87, 127.57, 126.56, 126.46, 76.34, 71.32, 64.55, 55.57, 32.70; MS 790 ($\text{M}+\text{NH}_4$, 100%).

Example 9F

10

By the procedure given above Example 9C, a solution of Example 9D (60mg, 0.078mmol), sodium cyanoborohydride (70mg, 1.1mmol), trace amounts of methyl orange in methanol was treated with 2N HCl (approx. 0.5 ml). Purification on T.L.C. plate with 40% ethyl acetate in methylene chloride gave the product (30mg). ^1H NMR (CDCl_3): δ 8.05 (s, 1H), 7.43- 7.06 (m, 28H), 5.13 (s, 2H), 4.90 (d, $J=14.3\text{Hz}$, 1H), 4.87 (d, $J=14.3\text{Hz}$, 1H), 4.61 (s, 2H), 3.98 (s, 2H), 3.59-3.49 (m, 4H), 3.07- 2.95 (m, 6H). ^{13}C NMR(CDCl_3): 161.98, 148.59, 139.48, 138.86, 138.32, 138.05, 137.74, 137.48, 132.56, 130.61, 129.87, 129.51, 128.91, 128.67, 128.65, 128.44, 128.37, 128.11, 127.91, 127.86, 127.66, 126.59, 126.56, 126.40, 77.20, 76.41, 76.15, 71.52, 71.48, 64.33, 64.11, 56.26, 55.58, 55.57, 33.79, 32.47. MS: 792 ($\text{M}+\text{NH}_4$, 40%) 775 ($\text{M}+\text{H}$, 100%).

Example 9G

To a suspension solution of bis (m-(HO)- $\text{C}_6\text{H}_4\text{CH}_2$) cyclic urea (280mg, 0.52 mmol) and Cs_2CO_3 (1.5g, 4.6mmol) in THF (4ml) was added methylchloroformate (122.9mg, 1.3mmol) and the resulting mixture was stirred at room temperature overnight. The mixture was filtered through celite and concentrated to give a residue which was purified on T.L.C. plate with 15% ethyl acetate in

methylene chloride to give the product (220 mg). as a solid, M.P. 146-147°C. ¹H NMR (CD₃OD): δ 7.34-6.95 (m, 18H), 4.78 (d, J=14.3Hz, 2H), 3.78 (s, 6H), 3.68 (bs, 2H), 3.61 (d, J=11.4Hz, 2H), 3.10- 2.83 (m, 6H). ¹³C NMR (CD₃OD): 162.22, 154.08, 151.28 139.74, 129.28, 129.14, 128.11, 126.52, 125.99, 121.63, 119.85, 70.44, 66.14, 55.46, 54.51, 32.22; MS: 655 (M+H, 100%).

Example 9I

10

To a solution of Example 9S (100mg) in tetrahydrofuran (1ml) was added methyl amine (0.4ml, 40% in water) and the resulting solution was stirred overnight. After concentration and purification on the plate, the product was obtained in good yield.

Examples 9J and 9K

By the procedure described previously for preparation of Example 9D, and substituting b-ethanolamine and trace amounts of molecular sieves (powder) in ethanol. The reaction mixture was filtered through Celite and concentrated to give a residue, which was purified on reverse phase T.L.C. plate with 90% methanol in water to the desired compounds.

Example 9J: ¹H NMR (CD₃OD): δ 8.31 (s, 2H), 7.67 (d, J=7.7Hz, 2H), 7.59 (s, 2H), 7.43- 7.03 (m, 14H), 4.74 (d, J=14Hz, 2H), 3.81-3.58 (m, 12H), 3.10- 2.86 (m, 6H); ¹³C NMR (CD₃OD): 164.97, 163.73, 141.22, 140.14, 137.66, 133.06, 130.65, 130.53, 130.06, 129.56, 128.50, 127.46, 71.93, 67.90, 64.26, 62.29, 57.20, 33.68, MS: 649 (M+H, 100%).

Example 9K: ¹H NMR (CD₃OD): δ 8.31 (s, 1H), 7.68- 7.03 (m, 18H), 5.33 (s, 1H), 4.76 (d, J= 13.9Hz, 1H), 4.75

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(d, $J=14.3\text{Hz}$, 1H), 3.81- 3.55 (m, 10H), 3.12- 2.7 (m, 8H); ^{13}C NMR (CD_3OD): 164.97, 163.78, 141.29, 141.24, 140.18, 139.39, 137.63, 133.06, 130.67, 130.64, 130.53, 130.48, 130.05, 129.55, 129.50, 128.86, 128.49, 127.47, 127.45, 127.20, 104.35, 71.98, 67.90, 67.40, 64.25, 62.39, 57.22, 57.11, 53.23, 33.67; M.S.: 649 (M+H, 100%).

Example 9L

10

To a solution of Example 5I (0.7g, 1.3 mmol) and triethylamine (0.263g, 2.6mmol) in THF (5ml) was added benzyl isocyanate (0.346g, 2.6mmol) and the resulting solution was stirred at room temperature overnight.

15 After removal of all the volatiles, the residue was purified on T.L.C. plate to give 0.7g of a solid, M.P. 150°C (decompose). ^1H NMR (CD_3OD): δ 7.33- 6.83 (m, 28H), 4.74 (d, $J=13.9\text{Hz}$, 2H), 4.30 (s, 4H), 3.64 (bs, 2H), 3.59 (d, $J=12.1\text{ Hz}$, 2H), 3.13- 2.91 (m, 6H); ^{13}C NMR (CD_3OD): 162.54, 155.68, 151.33, 139.88, 139.30, 138.64, 129.29, 129.20, 128.24, 128.18, 127.02, 126.90, 126.13, 125.81, 122.44, 120.55, 70.44, 65.28, 55.20, 44.31, 32.28; M.S: 805 (M+H, 100%).

25

Example 9M

To a solution of Example 5I (100mg, 0.186mmol) triethylamine (38mg. 0.39mmol) in THF (1 ml) was added methyl isocyanate (27mg. 0.47mmol) at room temperature and the resulting mixture was stirred overnight. After removal of all volatile reagents, a residue was purified on T.L.C. plate with 40% ethyl acetate in dichloromethane to give 51mgs of a solid, M.P. 150°C (decompose). ^1H NMR (CD_3OD) δ 7.44-6.99 (m, 18H), 4.82 (d, $J=14.2\text{Hz}$, 2H), 3.69- 3.65 (m, 4H), 3.15-2.95 (m, 6H), 2.84 (s, 6H). ^{13}C NMR (CD_3OD): 163.96, 157.56,

35

152.92, 141.31 140.83, 130.66, 130.55, 129.58, 127.46,
127.17, 123.70, 121.95, 71.88, 67.02, 56.67, 33.65,
27.57. M.S.: 670 (M+NH, 100%).

5

Example 9N

A solution of bis (m-bromobenzyl) cyclic urea
(MEM-protected) (0.84g, 1 mmol), propargyl alcohol
(0.224g, 4mmol), tetrakis (triphenylphosphine) palladium
10 (0.116g, 0.1mmol), copper iodide (0.019g, 0.1mmol) in
triethylamine (5ml) was refluxed under nitrogen
overnight. After evaporation of all volatiles, a
residue was diluted with ether (20ml) and filtered
through Celite. The filtrate was concentrated and
15 purified on T.L.C. plate to give 400mg of MEM-
protected-mono/coupling product. Deprotection of 170mg
of the coupled product by the standard procedure gave
130 mg of the desired product. ¹H NMR (CD₃OD): δ 7.50-
7.09 (m, 18H), 4.72 (d, J=13.9Hz, 1H), 4.69 (d,
20 J=13.4Hz, 1H), 4.46 (s, 2H), 3.75-3.68 (m, 4H), 3.18-
2.86 (m, 6H). ¹³C NMR (CD₃OD): 163.61, 141.99, 141.07,
139.73, 133.64, 133.54, 131.91, 131.76, 131.47, 130.57,
130.50, 129.81, 129.59, 129.19, 127.51, 124.69, 123.46,
89.28, 85.09, 71.93, 71.91, 68.12, 67.75, 57.07, 57.03,
25 51.15, 33.60, 33.59;; M.S.: 639/641 (M+H, 100%),
656/658 (M+NH₄, 100%).

Examples 9o and 9P

30

A solution of bis (m-Br-C₆H₄CH₂) cyclic urea
(425,mg, 0.64 mmol), 1-ethoxy-1-trimethylstannyl
ethylene (833mg, 3.84mmol) and Pd(PPh₃)₄ (37mg,
0.032mmol) in THF (5ml) was refluxed under N₂ overnight.
35 After cooling to room temperature, the reaction mixture
was diluted with ether (10ml) and filtered through

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silica gel to give two products. Further purification on T.L.C. plate with 20% ethyl acetate in methylene chloride gave Example 9Q (107mg, M.P. 190-191°C) and Example 9P (225 mg, M.P. 158-159°C).

5

Example 9o: ^1H NMR (CDCl_3): δ 7.78 (d, $J=7.3\text{Hz}$, 1H), 7.71 (s, 1H), 7.42- 7.02 (m, 16H), 4.81 (d, $J=13.9\text{Hz}$, 1H), 4.77 (d, $J=13.9\text{Hz}$, 1H), 3.71 (bs, 2H), 3.62-3.54 (m, 2H), 3.20- 2.85 (m, 8H), 2.50 (s, 3H); ^{13}C NMR (CDCl_3): 198.27, 161.86, 140.38, 139.29, 139.22, 138.81, 137.22, 133.99, 132.36, 130.65, 130.09, 129.36, 129.32, 128.99, 128.86, 128.66, 127.70, 127.55, 126.61, 122.53, 71.38, 65.24, 65.12, 55.83, 55.59, 32.77, 26.58, M.S.: 644/646 ($\text{M}+\text{NH}_4$, 100%).

15

Example 9P: ^1H NMR (CDCl_3): δ 7.78- 7.03 (m, 18H), 4.83 (d, $J=14.3\text{Hz}$, 2H), 3.73 (bs, 2H), 3.60 (d, $J=10.6\text{Hz}$, 2H), 3.17- 3.08 (m, 4H), 2.92-2.89 (m, 2H), 2.49 (s, 6H); ^{13}C NMR (CDCl_3): 198.03, 161.83, 139.40, 138.78, 137.12, 133.87, 129.29, 129.04, 128.74, 128.52, 127.36, 126.46, 71.25, 71.15, 65.32, 55.82, 30.73, 26.51; M.S.: 608 ($\text{M}+\text{NH}_4$, 100%).

25

Example 9Q

A solution of Example 9P (84mg, 0.142mmol) and hydroxylamine hydrochloride (59.4mg, 0.854mmol) in pyridine /ethanol (6ml, 1:1) was refluxed overnight. Evaporation of all solvents under vacuum gave a residue which was purified on preparative T.L.C. plates with ethyl acetate: methylene chloride: methanol (50:50:2) to give 71mg of a solid, M.P. 200-202°C. ^1H NMR (CD_3OD): δ 7.67- 7.07 (m, 18H), 4.74 (d, $J=13.9\text{Hz}$, 2H), 3.64-

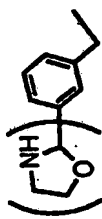
3.62 (m, 4H), 3.09- 2.89 (m, 6H), 2.17 (s, 6H); ^{13}C
NMR(CD₃OD): 163.94, 155.43, 141.23, 139.46, 138.97,
130.77, 130.65, 129.69, 129.37, 128.10, 127.47, 126.32,
72.02, 67.16, 57.08, 33.62, 11.96; M.S.: 621, (M+H,

5 100%).

Table 2d

Ex No.	Stereo	R ²²	R ²³	HPLC IC ₉₀ K ₁	mp, °C	MS	
						M+H	Notes
<u>2.3.4.5</u>							
9A	RSSR	m- (H0-N=CH) - C ₆ H ₄ CH ₂ -	m- (HONMCH ₂) - C ₆ H ₄ CH ₂ -	+++	214-216	595	(M+NH ₄)
9B	RSSR	m- (HONHCH ₂) - C ₆ H ₄ CH ₂ - HOAc	m- (HONHCH ₂) - C ₆ H ₄ CH ₂ - HOAc	+++		*	
9C	RSSR	m- (HONHCH ₂) - C ₆ H ₄ CH ₂ -	m- (HONHCH ₂) - C ₆ H ₄ CH ₂ -	+++		610	
9D	RSSR	m- (C ₆ H ₅ CH ₂ ON=CH) - C ₆ H ₄ CH ₂	m- (C ₆ H ₅ CH ₂ ON=CH) - C ₆ H ₄ CH ₂	++	170.5- 171	790	
9E	RSSR	m- (HONHCH ₂) - C ₆ H ₄ CH ₂ - HCl	m- (HONHCH ₂) - C ₆ H ₄ CH ₂ - HCl	+++		*	
9F	RSSR	m- (C ₆ H ₅ CH ₂ ON=CH) - C ₆ H ₄ CH ₂ -	m- (C ₆ H ₅ CH ₂ ON=CH) - C ₆ H ₄ CH ₂ -	++		775 (792)	
9G	RSSR	m- (CH ₃ OC (=O) -O) - C ₆ H ₄ CH ₂	m- (CH ₃ OC (=O) -O) - C ₆ H ₄ CH ₂	+++	146-147	655	
9H	RSSR	m- (CH ₃ CH (OH)) - C ₆ H ₄ CH ₂ -	m- (CH ₃ CH (OH)) - C ₆ H ₄ CH ₂ -			612	

9I	RSSR	m-(CH ₃ NHCH ₂)- C ₆ H ₄ CH ₂	m-(CH ₃ NHCH ₂)- C ₆ H ₄ CH ₂		593
9J	RSSR	m-(HOCH ₂ CH ₂ N=CH)- C ₆ H ₄ CH ₂ -	m-(HOCH ₂ CH ₂ N=CH)- C ₆ H ₄ CH ₂ -	+++ +++	649
9K	RSSR	m-(HOCH ₂ CH ₂ N=CH)- C ₆ H ₄ CH ₂ -	m-(HOCH ₂ CH ₂ N=CH)- C ₆ H ₄ CH ₂ -	+++ +++	649
9L	RSSR	m-(C ₆ H ₅ CH ₂ NHC(=O))- C ₆ H ₄ CH ₂ -	m-(C ₆ H ₅ CH ₂ NHC(=O))- O-C ₆ H ₄ CH ₂ -	+++ +++	805
9M	RSSR	m-(CH ₃ NHC(=O))- O-C ₆ H ₄ CH ₂ -	m-(CH ₃ NHC(=O))- O-C ₆ H ₄ CH ₂ -	+++ +++	670
9N	RSSR	m-(HOCH ₂ CC)- C ₆ H ₄ CH ₂ -	m-Br-C ₆ H ₄ CH ₂ -	+++ +++	639/641 (656/ 658)
9O	RSSR	m-(CH ₃ C(=O))- C ₆ H ₄ CH ₂ -	m-Br-C ₆ H ₄ CH ₂ -	+++ +++	190-191 644/646
9P	RSSR	m-(CH ₃ C(=O))- C ₆ H ₄ CH ₂ -	m-(CH ₃ C(=O))- C ₆ H ₄ CH ₂ -	+++ +++	158-159 608
9Q	RSSR	m-(CH ₃ C(=NOH))- C ₆ H ₄ CH ₂ -	m-(CH ₃ C(=NOH))- C ₆ H ₄ CH ₂ -	+++ +++	200-202 621
9R	RSSR	m-(CH ₃ CH(OH))- C ₆ H ₄ CH ₂ -	m-Br-C ₆ H ₄ CH ₂ -	+++ +++	629/631



9S	RSSR	m-(ClCH ₂)- C ₆ H ₄ CH ₂ -	m-(ClCH ₂)- C ₆ H ₄ CH ₂ -	+++	+++	156-157	603
9T	RSSR	m-(5-tetrazolyl)- -C ₆ H ₄ CH ₂ -	m-(5-tetrazolyl)- C ₆ H ₄ CH ₂ -	+++	+	159.4	643.2893
9U	RSSR	m-(5-tetrazolyl)- -C ₆ H ₄ CH ₂ -	cyclopropyl- methyl	+++	+++	171.6	539.2771
10A	RSSR	m-(CO ₂ H)- C ₆ H ₄ CH ₂ -	H	+++	+	141-143	2
10C	RSSR	m-(NC)- C ₆ H ₄ CH ₂ -	m-(OHC)-C ₆ H ₄ CH ₂ -	+++	+++	189-191	560
10D	RSSR	m-(CH ₃ ON=CH)- C ₆ H ₄ CH ₂ -	m-(CH ₃ ON=CH)- C ₆ H ₄ CH ₂ -	++	+	183-185	(577) 638
10E	RSSR	p-(CH ₃ C(=O)O)- C ₆ H ₄ CH ₂ -	p-(HO)-C ₆ H ₄ CH ₂ -	+++	+++	110-112	623
10F	RSSR	m-(CH ₃ C(=O)O)- C ₆ H ₄ CH ₂ -	m-(CH ₃ C(=O)O)- C ₆ H ₄ CH ₂ -	++	+++	623	8
10G	RSSR	m-(NH ₂ C(=O))- C ₆ H ₄ CH ₂ -	m-(NH ₂ C(=O))- C ₆ H ₄ CH ₂ -	+++	+++	(610)	7
10H	RSSR	m-(NH ₂ C(=O))- -CH ₂ O-C ₆ H ₄ CH ₂ -	m-(NH ₂ C(=O))- -CH ₂ O-C ₆ H ₄ CH ₂ -	+++	+++	NMR only	
10I	RSSR	m-(HO)-C ₆ H ₄ CH ₂ -	H	+++	+++	193-194	433
10J	RSSR	m-(CH ₃)-C ₆ H ₄ CH ₂ -	m-(HO)-C ₆ H ₄ CH ₂ -	+++	+++	537	3

10K	RSSR	2-Naphthylmethyl	m-(HO)-C ₆ H ₄ CH ₂ -	+++	+++	137-138	573	3
10L	RSSR	p-(CH ₃ C(=O)-O)- C ₆ H ₄ CH ₂ -	p-(CH ₃ C(=O)-O)- C ₆ H ₄ CH ₂ -	++	+++		623	8
10M	RSSR	m-(NH ₂ NHC(=O))- C ₆ H ₄ CH ₂ -	m-(NH ₂ NHC(=O))- C ₆ H ₄ CH ₂ -	+++	+++	205	623	7
10N	RSSR	p-(NH ₂ NHC(=O))- C ₆ H ₄ CH ₂ -	p-(NH ₂ NHC(=O))- C ₆ H ₄ CH ₂ -	+++	+++	215-217	623	7
10O	RSSR	p-(HOCH ₂)- C ₆ H ₄ CH ₂ -	m-(HO)-C ₆ H ₄ CH ₂ -	+++	++	141-144	553	3
10P	RSSR	m-(NH ₂ C(=O))- NHN=CH)-C ₆ H ₄ CH ₂ -	m-(NH ₂ C(=O))- NHN=CH)-C ₆ H ₄ CH ₂ -	+++	+++	301-303		5
10Q	RSSR	2-picolinyl-	m-(HO)-C ₆ H ₄ CH ₂ -	++	+++		524	3
10R	RSSR	m-(CH ₃ ONHC(=O))- C ₆ H ₄ CH ₂ -	m-(CH ₃ ONHC(=O))- C ₆ H ₄ CH ₂ -	+++	+++	150-158	(670)	7
10S	RSSR	p-(CH ₃ ONHC(=O))- C ₆ H ₄ CH ₂ -	p-(CH ₃ ONHC(=O))- C ₆ H ₄ CH ₂ -	+++	+++	186-189	653	7
10T	RSSR	m-(HOCH ₂ CH(OH)- CH ₂ O)-C ₆ H ₄ CH ₂ -	m-(HOCH ₂ CH(OH)- CH ₂ O)-C ₆ H ₄ CH ₂ -	+++	+++	113-115		9
10U	RSSR	m-(adamantamido)- C ₆ H ₄ CH ₂ -	m-(adamantamido)- C ₆ H ₄ CH ₂ -	+++	+++	183-184	893	8
10V	RSSR	m-(HO)-C ₆ H ₄ CH ₂ -	m-(adamantamido)- C ₆ H ₄ CH ₂ -	+	+++	196-198	716	6

10W	RSSR	m-(CH ₃ CH ₂ OC(=O))- -C ₆ H ₄ CH ₂ -	m-(CH ₃ CH ₂ OC(=O))	+++	+	178~180	651	10
10X	RSSR	m-(HONHC(=O))- C ₆ H ₄ CH ₂ -	m-(HONHC(=O))- C ₆ H ₄ CH ₂ -	+++	+++	139~143		11
10Y	RSSR	m-(HOCH ₂ CH ₂ O)- C ₆ H ₄ CH ₂ -	m-(HOCH ₂ CH ₂ O)- C ₆ H ₄ CH ₂ -	+++	+++	245~247	627	12
10Z	SRRS	p-(HOCH ₂)C ₆ H ₄ CH ₂	p-(HOCH ₂)C ₆ H ₄ CH ₂	+	++	198~199	567	1
11A	RSSR	m-(NH ₂ C(=NH))- C ₆ H ₄ CH ₂ -HOAC	m-(NH ₂ C(=NH))- C ₆ H ₄ CH ₂ -HOAC	++	+	224~226	591	13
11B	RSSR	(HOCH ₂ -CH(OH))- C ₆ H ₄ CH ₂ -	(HOCH ₂ -CH(OH))- C ₆ H ₄ CH ₂ -	+++	+++	135~137	627	9
11C	RSSR	m-(NH ₂ C(=O))- C ₆ H ₄ CH ₂ -	m, p, positions mixture	+++	+++	229~231		4
11D	RSSR	p-(HO)-C ₆ H ₄ CH ₂ -	p-(HO)-m-(HOCH ₂)- C ₆ H ₃ CH ₂ -	+++	++	178~180		14
11E	RSSR	p-(HO)-C ₆ H ₄ CH ₂ -	p-(HO)-m-(OHC)- C ₆ H ₃ CH ₂ -	+++	+++	567		14
11F	RSSR	p-(CH ₃ CH ₂ OC(=O))- C ₆ H ₄ CH ₂ -	p-(CH ₃ CH ₂ OC(=O))- C ₆ H ₄ CH ₂ -	++	+	174~178	651	10
11G	RSSR	m-(CH ₃ OC(=O))- C ₆ H ₄ CH ₂ -	m-(CH ₃ NHC(=O))- C ₆ H ₄ CH ₂ -	+++	+++	158~161	622 (639)	4
11H	RSSR	m-(CH ₃ NHC(=O))- C ₆ H ₄ CH ₂ -	m-(CH ₃ NHC(=O))- C ₆ H ₄ CH ₂ -	+++	+++	160~163	621	7

11I	RSSR	benzyl	m-(HO)-C ₆ H ₄ CH ₂ -	+++	+++	189-193	523	3
11J	RSSR	2-Naphthylmethyl	p-(HOCH ₂)-C ₆ H ₄ CH ₂	+++	+++	199-201	587	3
11K	RSSR	m-(HOCH ₂ -CH(OH))	m-(HOCH ₂ -CH(OH))	+++	+++		627	9
		-C ₆ H ₄ CH ₂ -	-C ₆ H ₄ CH ₂ -					
11L	RSSR	p-(HOCH ₂ -CH(OH))	p-(HOCH ₂ -CH(OH))	+++	+++		627	9
		-C ₆ H ₄ CH ₂ -	-C ₆ H ₄ CH ₂ -					
11M	RSSR	m-(HOCH ₂)	m-(CH ₃ NHC(=O))-	+++	+++	107-109	(611)	4
		-C ₆ H ₄ CH ₂ -	C ₆ H ₄ CH ₂ -				594	
11N	RSSR	p-(HOCH ₂)-	H	+++	+++	112-114	447	2
		C ₆ H ₄ CH ₂ -						
11O	RSSR	m-((HO) ₂ B)-	m-((HO) ₂ B)-	+++	+++	263-267		15
		C ₆ H ₄ CH ₂ -	C ₆ H ₄ CH ₂ -					
11P	RSSR	m-(NO ₂)-C ₆ H ₄ CH ₂ -	m-(C ₆ H ₅ CH ₂ O)-	+	+++	173-176	(675)	3
			C ₆ H ₄ CH ₂ -					
11Q	RSSR	m-(NH ₂)-C ₆ H ₄ CH ₂ -	m-(HO)-C ₆ H ₄ CH ₂	+++	+++	116-120	538	3
11R	RSSR	m-(CH ₃ CH ₂ NHC(=O))-C ₆ H ₄ CH ₂ -	m-(CH ₃ CH ₂ NHC(=O))-	+++	+++	135-138	649	7
			C ₆ H ₄ CH ₂ -					
11S	RSSR	m-((CH ₃) ₂ N	m-((CH ₃) ₂ NC(=O))-	+++	+++	132-134	649	7
		C(=O))-C ₆ H ₄ CH ₂ -	C ₆ H ₄ CH ₂ -					
11T	RSSR	m-(CH ₃ O ₂ C)-	m-((CH ₃ CH ₂) ₂ NC				(681)	4
		C ₆ H ₄ CH ₂	(=O))-C ₆ H ₄ CH ₂ -	+++	+++			
11U	RSSR	m-(CH ₃ O ₂ C)-	m-(CH ₃ CH ₂ NHC	+++	+++		(653)	4
		C ₆ H ₄ CH ₂ -	(=O))-C ₆ H ₄ CH ₂ -					
12A	RSSR	6-amino-1-hexyl	6-amino-1-hexyl	+	+		525	

12B	RSSR	6-amino-1-hexyl	benzyl	++	+	516
12C	RSSR	6-hydroxy-1-hexyl	H	+++	+++	427
12D	RSSR	hexyl	6-hydroxy-1-hexyl	+++	+++	527
12E	RSSR	hexyl	6-hydroxy-1-hexyl	+++	+++	481
12F	RSSR	cyclopropyl	hexyl	++	++	471
12G	RSSR	methyl	4-hydroxy-1-butyl	+++	+++	555
12H	RSSR	butyl	5-carboxy-1-pentyl	++	++	583
12I	RSSR	5-carboxy-1-pentyl	5-carbomethoxy-1-pentyl	+++	+	759
12J	RSSR	1-pentyl	5-carboxy-1-pentyl	+++	+++	583
12K	RSSR	3-iodobenzyl	8-hydroxy-1-octyl	++		495
12L	RSSR	8-hydroxy-1-octyl	2-(hydroxy methyl)-cyclopropyl-methyl	+++	+++	501
12M	RSSR	benzyl	2-(hydroxy methyl)-cyclopropyl-methyl	++	+	411
12N	RSSR	2-(hydroxy methyl)-cyclopropyl-methyl	H	++	++	411
12O	RSSR	7-hydroxy-1-heptyl	7-hydroxy-1-heptyl	++	+++	555
12P	RSSR	7-hydroxy-1-heptyl	3-(carbamoyl thio)benzyl	+++	+	655
12Q	RSSR	3-(carbamoyl thio)benzyl	(3-methylthio)benzyl	+++	+++	599
12R	RSSR	(3-methylthio)benzyl				

12S	RSSR	3- (methylsulfonyl) benzyl	3- (methylsulfonyl) benzyl	+++	+++	633
12T	RSSR	2-[(2-hydroxy ethyl)oxy]ethyl	H	++	+	415
12U	RSSR	2-[(2-hydroxy ethyl)oxy]ethyl	2-[(2-hydroxy ethyl)oxy]ethyl	+	++	503
12V	RSSR	6-acetoxy-1- hexyl	6-hydroxy-1- hexyl	++	+++	569
12W	RSSR	6-(N-methylamino carboxy)-1-hexyl	6-(N-methylamino carboxy)-1-hexyl	++	++	641
12X	RSSR	6-(N-methylamino carboxy)-1-hexyl	6-acetoxy-1- hexyl	++	+++	626
12Y	RSSR	6-(N-methylamino carboxy)-1-hexyl	6-acetoxy-1- hexyl	+++		584
12Z	RSSR	2-(2-hydroxy ethyl) cyclopropyl- methyl	H	++		425
						16
13A	RSSR	2-(3- hydroxypropyl)- cyclopropyl- methyl	H	++		439
						16
13B	RSSR	[2-(2- hydroxypropyl)- 3,3-dimethyl] cyclopropyl methyl	H	++		481
						16
13C	RSSR	[2-(2- hydroxypropyl)- 3,3-dimethyl] cyclopropyl- methyl	[2-(2- hydroxypropyl)- 3,3-dimethyl] cyclopropyl- methyl	+		635 (652)
13D	RSSR	6-hexenyl	6-hexenyl	+++	+++	491
13E	RSSR	5,6-epoxy-1- hexyl	5,6-epoxy-1- hexyl	++	+++	523
13F	RSSR	5,6-dibromo-1- hexyl	5,6-dibromo-1- hexyl	++	+	811

13G	RSSR	6-bromo-5-hydroxy-1-hexyl	6-bromo-5-hydroxy-1-hexyl	+++	+++	685
13H	RSSR	5-hydroxy-1-pentyl	5-hydroxy-1-pentyl	+++	+++	499
13I	RSSR	5,6-dihydroxy-1-hexyl	5,6-dihydroxy-1-hexyl	+++	+	559
13J	RRSR	cyclopropyl-methyl	cyclopropyl-methyl	+++	+++	435
13K	RRSR	allyl	allyl	+++	+++	407
13L	RRSR	benzyl	benzyl	+++	+++	507
13M	RRSR	4-(hydroxy-methyl)benzyl	4-(hydroxy-methyl)benzyl	+++	+++	567
13N	RRRR	allyl	allyl	+	++	407
13O	RRRR	cyclopropyl-methyl	cyclopropyl-methyl	+	++	435
13P	RRRR	H	H	+	+	327
13Q	RRRR	benzyl	benzyl	+	+++	507
13R	RRRR	n-butyl	n-butyl	++	+++	439
13S	RRRR	4-(hydroxy-methyl)benzyl	4-(hydroxy-methyl)benzyl	++	++	567
13T	RSSR	(1,2,3,4-tetrahydro)-6-isoquinolyl-methyl	(1,2,3,4-tetrahydro)-6-isoquinolyl-methyl	++	+	617

13U	RSSR	6-isoquinolyl- methyl	6-isoquinolyl- methyl	++	+	264-6	609
13V	RSSR	2-thiazolyl- methyl	2-thiazolyl- methyl	++	++		521
13W	RSSR	(5-t- butoxycarbonyl)- 2-furanylmethyl	(5-t- butoxycarbonyl)- 2-furanylmethyl	+	+++	182-4	687
13X	RSSR	(5- hydroxymethyl)- 2-furanylmethyl	(5- hydroxymethyl)- 2-furanylmethyl	++	+		547
13Y	RRRS	benzyl	benzyl	+	+++		507
13Z	RSSR	4-chloro-3- pyridylmethyl	4-chloro-3- pyridylmethyl	++	+++		577 (594)
14A	RSSR	2-chloro-3- pyridylmethyl	2-chloro-3- pyridylmethyl	++	++		685
14B	RSSR	2-(t-butylthio)- 3-pyridylmethyl	2-(t-butylthio)- 3-pyridylmethyl	++	+++		541
14C	RSSR	4-hydroxy-3- pyridylmethyl	4-hydroxy-3- pyridylmethyl	++	+		458
14D	RSSR	cyclopropyl- methyl	2-pyridyl	++	+++		543 739
14E	RSSR	oct-2-yn-1-yl	oct-2-yn-1-yl	++			16
14F	RSSR	3,3-diphenyl- 2(R)-	3,3-diphenyl- 2(R)-	++			479 587
14G	RSSR	cyclopropylmethyl	cyclopropylmethyl	++			16
14H	RSSR	phenyl	phenyl	++			457
		3(S)-phenyl- 2(R)-	3(S)-phenyl- 2(R)-	++			
14I	RSSR	cyclopropyl- methyl	cyclopropyl- methyl	++			
		3(S)-phenyl- 2(R)-	H	++			
14J	RSSR	cyclopropyl- methyl	3-benzoyloxy-5- methyl-4- pyridylmethyl	++			
14K	RSSR	3-benzoyloxy-5- methyl-4- pyridylmethyl	3-benzoyloxy-5- methyl-4- pyridylmethyl	++	+		
		3-hydroxy-5- methyl-4- pyridylmethyl	3-hydroxy-5- methyl-4- pyridylmethyl	++			

15A	RSSR	m-(H ₂ N)-C ₆ H ₄ CH ₂ -	p-(HOCH ₂)- C ₆ H ₄ CH ₂ -	+++	+++	552.29
15B	RSSR	m-(CF ₃ O ₂ SNH)- C ₆ H ₄ CH ₂ -	m-(CF ₃ O ₂ SNH)- C ₆ H ₄ CH ₂ -	+++	++	801.19
15C	RSSR	m-(H ₂ NCONH)- C ₆ H ₄ CH ₂ -	m-(H ₂ NCONH)- C ₆ H ₄ CH ₂ -	+++	+++	565.32
15D	RSSR	m-(CH ₃ NH)- C ₆ H ₄ CH ₂ -	m-(CH ₃ NH)- C ₆ H ₄ CH ₂ -	+++	+++	822.39
15E	RSSR	m-(C ₆ H ₅ CH ₂ OCONH)- C ₆ H ₄ CH ₂ -	m-(C ₆ H ₅ CH ₂ OCONH)- C ₆ H ₄ CH ₂ -	++	+	190.7
15F	RSSR	m-(CH ₃ (OHC)N)- C ₆ H ₄ CH ₂ -	m-(CH ₃ (OHC)N)- C ₆ H ₄ CH ₂ -	++	+++	621.31
15G	RSSR	cyclopropyl- methyl	m-(H ₂ N)-C ₆ H ₄ CH ₂ - HCl	+++	+++	486.27
15H	RSSR	cyclopropyl- methyl	m-(O ₂ N)-C ₆ H ₄ CH ₂ -	+++	+++	516.25
15I	RSSR	m-(CH ₃) ₂ N)- C ₆ H ₄ CH ₂ -	m-(CH ₃) ₂ N)- C ₆ H ₄ CH ₂ -	+++	+++	593.35
15J	RSSR	m-(O ₂ N)-C ₆ H ₄ CH ₂ -	p-(HOCH ₂)- C ₆ H ₄ CH ₂ -	+++	+++	(599)
15K	RSSR	m-(H ₂ N)-C ₆ H ₄ CH ₂ -	m-(CH ₃ NHCONH)- C ₆ H ₄ CH ₂ -	+++	+++	594.5
15L	RSSR	m-(CH ₃ NHCONH)- C ₆ H ₄ CH ₂ -	m-(CH ₃ NHCONH)- C ₆ H ₄ CH ₂ -	+++	+++	(651.4)

Notes (for Table 2d):

- (1) Prepared according to the general alkylation procedure.
- 5 (2) Monoalkyl compounds were prepared by following the procedure 5 under the title of synthesis of monoalkyl cyclic urea.
- 10 (3) Prepared by the alkylating the appropriate monoalkyl compound.
- (4) Isolated as the side product due to the uncompletion reaction.
- 15 (5) Hydroxylamine hydrochloride (0.384 mmol) was added into appropriate bis(N-m-benzaldehyde) cyclic urea (0.064 mmol) in a mixture of 2mL pyridine and 2mL ethanol. The mixture was heated to reflux for 4
- 20 hr and the solvent was removed on a rotary evaporator. The residue was purified on silica gel (0.2:3:7 methanol:ethyl acetate:methylene chloride).
- 25 (6) Preparation of monoalkyl compounds were favored by using one equivalent of alkylating agent.
- (7) A solution of bis(N-m-benzoic acid) cyclic urea (1.56 mmol) in 1:1 benzene: methylene chloride
- 30 containing 1 drop of DMF and pyridine (9.36 mmol) was treated with oxalyl chloride (2M in methylene chloride, 9.36 mmol) at zero degree and stirred at room temperature for overnight. The solvent removed
- 35 on a rotary evaporator and the resulting residue was dried under pump for 2 hr. To the residue 20 mL methylene chloride was added followed by pyridine

(9.36 mmol) and methylamine (8.03M in ethanol, 9.36 mmol). Stirred at room temperature for overnight. The mixture was extracted with EtOAc, dried over MgSO₄ and purified on silica gel (1:9 methanol
5 chloroform).

(8) A solution of bis(N-m-hydroxy-benzyl)cyclic urea (0.93 mmol) in methylene chloride was treated with triethylamine (1.67 mmol) at -20°C. Then a
10 solution of 1-adamantyl isocyanate (1.4 mmol) in 5 mL methylene chloride was added dropwise. The mixture was stirred at -20°C for 10 min, 0°C for 1 hr, room temperature for overnight and washed with cooled 5% HCl, sat'd NaHCO₃, water, and dried over
15 MgSO₄. The residue was purified on silica gel (2:8 ethyl acetate: methylene chloride).

(9) 2 drops of OsO₄ (25% in t-BuOH) was added into the mixture of bis(N-m-allyloxy-benzyl) cyclic urea
20 (0.19 mmol) and N-methyl-morpholine N-oxide (0.57 mmol) in 5 mL acetone. The mixture was stirred at RT for overnight. The solvent was removed on a rotary evaporator and the residue was purified on silica gel (2:8 methanol:chloroform).

25

(10) The ethyl ester was prepared by refluxing appropriate acid with 4M HCl (in dioxane) in ethanol.

(11) To the appropriate ethyl ester (0.21 mmol) and hydroxylamine hydrochloride (1.71 mmol) mixture (10 mL methanol was used as the solvent), a 5M (2.1 mmol) solution of potassium hydroxide in methanol was added dropwisely at room temperature. After
30 stirring for 24 hr, methanol was evaporated and the solid residue acidified with acetic acid and

extracted with ethyl acetate which was purified on silica gel (0.1:3:7 acetic acid:methanol:ethyl acetate).

5 (12) To the bis(N-m-allyloxy-benzyl) cyclic urea (0.16 mmol) in 10mL methanol, ozone was bubbled through for 10 min at -78°C. After warmed up to RT, sodium borohydride (1.6 mmol) was added and stirred at RT for overnight. The reaction was worked up by
10 quenching with acetic acid, the solvent was removed on rotary evaporator and purified on silica gel (0.3:9.7 methanol:ethyl acetate).

(13) A solution of the bis-cyano compound in
15 absolute methanol was saturated with hydrochloric acid and left at zero degree for 3 hr followed by TLC until completed. After evaporation of the solvent, the resulting was solubilized in 2M ammonia in methanol. After one week stirring at
20 room temperature, the solvent was removed under reduced pressure and the residue was purified on silica gel (by 0.2:1:9 acetic acid:methanol:ethyl acetate) to give a solid.

25 (14) To the bis(N-p-hydroxy-benzyl) cyclic urea (1.11 mmol) in ethanol, NaOH (15.54 mmol) in 2 mL water was added dropwise. The mixture was heated up to 80°C for 1 hr, then 2 mL chloroform was added following additional heating at 80°C for overnight.
30 The reaction was worked up by neutralizing with 5% cooled HCl, washed with sat'd sodium bicarbonate, water and dried over MgSO₄. The residue was purified on silica gel (1.5:8.5 methanol:chloroform). To this solid, ethanol was
35 added following by excess sodium borohydride. The mixture was heated up to reflux for 2 hr. The

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reaction was worked up by quenching with acetic acid, the solvent was removed on rotary evaporator and the purified on silica gel (2:8 methanol:chloroform).

- 5
10
15
- (15) n-Butyllithium (1.6M in hexane, 5 mmol) was added dropwise to bis(N-m-bromo-benzyl) cyclic urea (1 mmol) in THF at -78°C. After stirring for 0.5 hr, trimethyl borate (5 mmol) was added. The mixture slowly warmed up to room temperature and remained stirring for 4 hr. The reaction mixture was decomposed by the addition of 5% HCl, diluted with ethyl acetate, washed 2 time with water and dried over MgSO₄. The residue was purified on silica gel (0.1: 0.5:9.5 acetic acid:methanol:chloroform).

- 20
- (16) Required alkylating agents were prepared in a three step sequence: 1.) Asymmetric cyclopropanation [D. A. Evans et al. , J. Amer. Chem. Soc. 1991 , 113 , 726-728]. 2.) Reduction with lithium aluminum hydride. 3.) Conversion to the bromide with CBr₄ , Ph₃P, and imidazole.

- 25
- Listed below are a representative list of data for compounds listed in Table 2d:

30

Example 12A: MS: 525 (M + 1). NMR (CD₃OD): δ 7.30-7.10 (m, 10H), 3.96-3.88 (m, 2H), 3.58-3.42 (m, 4H), 3.37-3.28 (m, 4H), 3.17-3.10 (m, 2H), 2.94-2.78 (m, 6H), 2.20-2.08 (m, 2H), 1.72-1.10 (br m, 14H).

35

Example 12B: MS: 516 (M + 1, 100). NMR (CD₃OD): δ 6.96-7.72 (br m, 15H), 4.68 (d, 1H), 3.78-3.89 (m, 1H), 3.43-3.64 (m, 3H), 3.28-3.34 (m, 2H),

3.10-3.19 (m, 1H), 2.81-3.02 (m, 6H), 2.12-2.26 (m, 1H), 1.10-1.84 (br m, 9H).

5 Example 12C: MS: 427 (M + 1, 100). NMR (CDCl₃):
δ 7.16-7.36 (m, 10H), 4.82 (d, 1H), 3.88-3.98 (m, 2H), 3.37-3.69 (m, 8H), 3.04-3.19 (m, 3H), 2.63-2.77 (m, 1H), 2.08-2.22 (m, 1H), 1.74-1.88 (m, 2H), 1.10-1.64 (br m, 6H).

10 Example 12D: MS: 527 (M + 1, 100). NMR (CDCl₃):
δ 7.17-7.38 (m, 10H), 3.99 (s, 2H), 3.45-3.75 (m, 8H), 3.03-3.14 (m, 2H), 2.89-3.01 (m, 2H), 2.60-2.68 (m, 2H), 2.12-2.24 (m, 2H), 1.12-1.62 (br m, 18H).

15 Example 12E: MS: 481 (M + 1, 100). NMR (CDCl₃):
δ 7.17-7.37 (m, 10H), 3.94-4.13 (m, 2H), 3.61-3.75 (m, 2H), 3.43-3.61 (m, 4H), 3.05-3.20 (m, 3H), 2.73-3.03 (m, 3H), 2.10-2.23 (m, 1H), 1.98-2.08 (m, 1H), 1.02-1.75 (br m, 9H), 0.82-0.97 (m, 1H), 0.30-0.47 (m, 2H), 0.02-0.13 (m, 2H).

25 Example 12F: MS: 471 (M + 1, 100). NMR (DMSO-d₆): δ 7.06-7.33 (m, 10H), 5.25 (s, 2H), 4.34 (t, 2H), 3.71 (s, 2H), 3.44-3.58 (m, 2H), 3.20-3.42 (m, 6H), 2.96-3.06 (m, 2H), 2.79-2.91 (m, 2H), 1.83-1.97 (m, 2H), 1.08-1.42 (br m, 8H).

30 Example 12G: FABMS: 555 (M + 1, 100). NMR (DMSO-d₆): δ 8.41 (s, 4H), 7.06-7.33 (m, 10H), 3.71 (s, 2H), 3.33-3.50 (m, 5H), 3.00-3.09 (m, 2H), 2.75-2.88 (m, 2H), 2.11-2.21 (m, 2H), 2.03-2.11 (m, 3H), 1.90-2.03 (m, 2H), 1.02-1.54 (br m, 10 H).

35 Example 12H: MS: 583 (M + 1, 100). NMR (CDCl₃):
δ 7.12-7.35 (m, 10H), 3.98 (s, 2H), 3.61 (s, 6H),

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3.60-3.74 (m, 2H), 3.45-3.58 (m, 2H), 3.03-3.13 (m, 2H), 2.84-3.03 (m, 4H), 2.07-2.28 (m, 6H), 1.08-1.81 (br m, 12H).

5 Example 12I: MS: 555 (M - CH₂ + 1, 100). NMR (CDCl₃): δ 7.12-7.34 (m, 10H), 3.98 (s, 2H), 3.62 (s, 3H), 3.45-3.76 (m, 6H), 3.04-3.13 (m, 2H), 2.85-2.99 (m, 3H), 2.79 (s, 1H), 2.10-2.28 (m, 4H), 1.08-1.64 (br m, 12H).

10 Example 12J: MS: 759 (M + 1, 100), 633 (M - I + 1, 33), 507 (M - 2I + 1, 17). NMR (CDCl₃): δ 7.00-7.64 (br m, 18H), 4.77 (d, 2H), 3.70 (s, 2H), 3.49-3.57 (m, 2H), 2.99-3.12 (m, 4H), 2.81-2.93 (m, 2H),
15 2.29 (s, 2H).

Example 12K: MS: 583 (M + 1, 100). NMR (CDCl₃): δ 7.15-7.35 (m, 10H), 3.98 (s, 2H), 3.44-3.74 (m, 8H), 3.02-3.22 (m, 2H), 2.88-3.02 (m, 2H), 2.02-2.20 (m, 2H), 0.94-1.92 (br m, 28H).

25 Example 12L: MS: 495 (M+1,100) NMR (CDCl₃): δ 7.31 (m,10H) 4.22 (m. 4H) 3.76-4.05 (m. 6H) 3.60 (m, 2H) 3.03-3.37 (m, 6H) 1.89 (m, 2H) 0.95 (m, 2H) 0.80 (m, 2H) 0.45 (m, 2H) 0.30 (m, 2H).

30 Example 12M: MS: 501 (M+1,100) NMR (CDCl₃): δ 7.05-7.38 (m, 15H) 4.84 (dd,1H) 4.04 (m, 1H) 3.93 (m, 1H) 3.49-3.75 (m. 3H) 3.05 (m, 6H) 1.74 (m, 2H) 0.70-0.98 (m, 2H) 0.40 (m, 1H) 0.28 (m, 1H).

Example 12N: MS: 411 (M+1, 100) NMR (CDCl₃): δ 7.25 (m, 10H) 4.72 (bs, 1H) 3.95 (m, 5H) 3.53 (m. 2H) 3.11 (m. 6H) 2.94 (t, 1H) 2.76 (m. 1H) 0.84 (m. 2H) 0.39 (m, 1H) 0.24 (m. 1H).

Example 12o: MS: 441 (M + 1, 100). NMR
(CDCl₃): δ 7.17-7.36 (m, 10H), 4.70 (d, 1H), 3.90-
4.01 (m, 2H), 3.53-3.68 (m, 4H), 3.37-3.45 (m, 1H),
3.04-3.20 (m, 3H), 2.78-3.03 (m, 1H), 2.68-2.77 (m,
5 1H), 2.09-2.24 (m, 1H), 1.10-1.95 (br m, 12H).

Example 12P: MS: 555 (M + 1, 100). NMR (CDCl₃):
 δ 7.13-7.37 (m, 10H), 3.98 (s, 2H), 3.44-3.77 (m,
10H), 3.03-3.14 (m, 2H), 2.80-3.01 (m, 4H), 2.06-
10 2.21 (m, 2H), 1.00-1.75 (br m, 20H).

Example 12Q: m. p. 185.7 ° C. FABMS: 655 (M + 1,
100), 613 (M - HNCNH + 1, 80). NMR (DMSO-d₆): δ
8.99 (br s, 8H), 6.87-7.62 (br m, 18H), 5.08-5.35
15 (m, 2H), 4.44-4.62 (m, 2H), 3.41-3.61 (m, 4H),
2.96-3.18 (m, 4H), 2.64-2.81 (m, 2H). Reference:
K. Takagi, Chemistry Letters (1985), pp. 1307-1308.

Example 12R: m. p. 169.3° C. MS: 599 (M + 1,
20 100). NMR (CDCl₃): δ 6.90-7.40 (m, 18H), 4.88 (d,
2H), 3.63 (s, 2H), 3.49-3.58 (m, 2H), 2.91-3.12 (m,
6H), 2.42 (s, 6H), 2.22 (s, 2H).

Example 12S: MS: 663 (M + 1), 680 (M + NH₃ + 1).
25 NMR (CDCl₃): δ 6.93-7.83 (br m, 18H), 4.78 (d, 2H),
3.71 (s, 2H), 3.53-3.59 (m, 2H), 3.19 (d, 2H),
3.00-3.14 (m, 4H), 2.98 (s, 6H), 2.75-2.83 (m, 2H).

Example 12T: MS: 415 (M + 1). NMR (Acetone-d₆):
30 δ 7.15-7.30 (m, 10H), 5.29 (s, 1H), 4.41 (s, 1H),
4.25 (s, 1H), 3.81-4.05 (m, 3H), 3.63-3.78 (m, 2H),
3.56-3.63 (m, 2H), 3.37-3.47 (m, 5H), 3.05-3.25 (m,
3H), 2.79-2.94 (m, 1H), 2.20 (br s, 1H).

35 Example 12U: MS: 503 (M + 1). NMR (Acetone-d₆):
 δ 7.15-7.28 (m, 10H), 4.29 (s, 1H), 3.95-4.07 (m,

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4H), 3.49-3.66 (m, 8H), 3.35-3.49 (m, 8H), 3.14-3.22 (m, 2H), 3.02-3.09 (m, 2H), 2.08-2.21 (m, 3H).

5 Example 12V: MS: 569 (M + 1). NMR (CDCl₃): δ 7.14-7.34 (m, 10H), 3.90-4.03 (m, 4H), 3.42-3.74 (m, 8H), 3.03-3.14 (m, 2H), 2.86-3.01 (m, 2H), 2.68-2.86 (m, 2H), 2.05-2.23 (m, 2H), 2.01 (s, 3H), 0.98-1.75 (br m, 15 H).

10 Example 12W: MS: 641 (M + 1), 584 (M - CONHCH₂ + 1), 527 (M - 2CONHCH₂ + 1). NMR (CDCl₃): δ 7.10-7.35 (m, 10H), 4.77 (br s, 2H), 3.89-4.12 (m, 6H), 3.60-3.80 (m, 2H), 3.35-3.47 (m, 2H), 3.04-3.16 (m, 2H), 2.87-3.01 (m, 2H), 2.78 (d, 6H), 2.01-2.16 (m, 2H), 0.98-1.80 (br m, 18H).

20 Example 12X: MS: 626 (M + 1), 569 (M - CONHCH₂ + 1). NMR (CDCl₃): δ 7.09-7.35 (m, 10H), 4.79 (br s, 1H), 3.82-4.14 (m, 6H), 3.38-3.81 (m, 4H), 3.01-3.22 (m, 2H), 2.83-3.01 (m, 2H), 2.79 (d, 3H), 2.04-2.20 (m, 2H), 2.01 (s, 3H), 0.96-1.84 (br m, 18 H).

25 Example 12Y: MS: 584 (M + 1), 527 (M - CONHCH₂ + 1). NMR (CDCl₃): δ 7.11-7.41 (m, 10H), 4.81 (br s, 1H), 3.87-4.10 (m, 4H), 3.41-3.76 (m, 6H), 3.02-3.16 (m, 2H), 2.81-3.02 (m, 2H), 2.78 (d, 3H), 2.00-2.21 (m, 2H), 1.79 (br s, 1H), 0.95-1.63 (br m, 18H).

30 Example 12Z: MS: 425 (M+1, 100).

Example 13A: MS: 439 (M+1, 100).

35 Example 13B: MS: 481 (M+1, 100).

Example 13C: MS: 635 (M+1, 100) 652 (M+NH₄).

Example 13D: MS: 491 (M+1,100); NMR (CDCl₃): δ
7.24 (m,10H), 5.68 (m,2H), 4.90 (m,4H), 3.98
5 (s,2H), 3.66 (m,2H), 3.50 (m,2H), 3.04 (m,4H), 2.19
(m,2H), 1.95 (m,4H), 1.22 (m,8H).

Example 13E: MS: 523 (M+1,100); NMR (CDCl₃): δ
7.19 (m,10H), 3.98 (s,2H), 3.66 (m,2H), 3.53
10 (m,2H), 3.23 (s,2H), 3.14-2.66 (m,8H), 2.40 (m,2H),
2.21 (m,2H), 1.55-1.23 (m,12H).

Example 13F: MS: 811 (M+1,100); NMR (CDCl₃): δ
7.22 (m,10H), 4.00 (m,4H), 3.77 (dd,2H), 3.56
15 (m,6H), 3.13 (m,4H), 2.93 (m,2H), 2.23 (m,2H), 2.00
(m,2H), 1.65 (m,2H), 1.50-1.23 (m,8H).

Example 13G: MS: 685 (M+1,100); NMR (CDCl₃): δ
7.22 (m,10H), 4.01 (m,2H), 3.72-3.26 (m,12H), 3.14
20 (d,2H), 2.96-2.70 (m,6H), 2.26 (m,2H), 1.43-1.18
(m,8H).

Example 13H: MS: 499 (M+1,100); NMR (CDCl₃): δ
7.23 (m,10H), 3.99 (s,2H), 3.54 (m,8H), 3.12
25 (m,2H), 2.94 (m,4H), 2.22 (m,2H), 1.47-1.20
(m,12H).

Example 13I: MS: 559 (M+1,100); NMR
(CDCl₃/CD₃OD): δ 7.02 (m,10H), 3.79 (s,2H), 3.69
30 (s,2H), 3.28 (m,6H), 3.13 (m,4H), 2.86 (m,2H), 2.67
(m,2H), 1.95 (m,2H), 1.09-0.90 (m,12H).

Example 13J: MS: 435 (M+1,100); NMR (CDCl₃): δ
7.26 (m,10H), 4.15 (m,1H), 3.86 (m,1H), 3.76
35 (m,1H), 3.64 (m,1H), 3.53 (m,2H), 3.23-2.97 (m,4H),
2.78 (m,2H), 2.16 (d,1H), 1.92 (m,1H), 1.18 (m,1H),

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0.88 (m,1H), 0.58 (m,2H), 0.46 (m,2H), 0.29 (d,1H),
0.07 (d,1H).

- Example 13K: MS: 407 (M+1,100); NMR (CDCl₃): δ
 5 7.23 (m,10H), 6.05 (m,1H), 5.67 (m,1H), 5.25
 (m,2H), 5.05 (dd,2H), 4.17 (m,2H), 3.96 (m,1H),
 3.83 (m,1H), 3.65 (m,1H), 3.55 (m,1H), 3.12-2.79
 (m,5H), 2.59 (br s,1H), 2.35 (d,1H).
- 10 Example 13L: MS: 507 (M+1,100); NMR (CDCl₃): δ
 7.28 (m,18H), 6.91 (d,2H), 4.10 (d,2H), 3.92
 (d,1H), 3.58 (m,4H), 3.18-2.63 (m,5H).
- Example 13M: MS: 567 (M+1,100); NMR (CDCl₃): δ
 15 7.46-6.95 (m,18H), 4.90 (dd,2H), 4.67 (br d,4H),
 3.84 (d,1H), 3.53 (m,4H), 3.09-2.75 (m,5H), 2.44
 (d,1H).
- Example 13N: MS: 407 (M+1,100); NMR (CDCl₃): δ
 20 7.24 (m,10H), 5.55 (m,2H), 5.14 (m,4H), 4.07
 (dd,2H), 3.74 (m,2H), 3.27 (m,2H), 3.12-2.84
 (m,6H), 2.09 (s,2H).
- Example 13o: MS: 435 (M+1,100); NMR (CDCl₃): δ
 25 7.26 (m,10H), 3.85 (s,2H), 3.41 (m,4H), 3.01
 (ddd,4H), 2.28 (dd,2H), 2.19 (s,2H), 0.93 (m,2H),
 0.50 (m,2H), 0.16 (m,2H).
- Example 13P: MS:327 (M+1,100); NMR (CDCl₃): δ
 30 7.28 (m,10H), 4.25 (m,2H), 3.35 (m,6H), 2.81
 (m,2H), 2.65 (m,2H).
- Example 13Q: MS: 507 (M+1,100); NMR (CDCl₃): δ
 35 7.24 (m,20H), 4.74 (d,2H), 3.55 (m,4H), 3.20
 (m,2H), 2.96 (m,2H), 2.77 (m,2H).

- Example 13R: MS: 439 (M+1,100); NMR (CDCl₃): δ 7.26 (m,10H), 3.75 (s,2H), 3.62 (m,2H), 3.20 (m,2H), 3.08 (m,2H), 2.84 (m,2H), 2.38 (m,2H), 1.38-1.23 (m,8H), 0.90 (m,6H).
- 5 Example 13S: MS: 567 (M+1,100); NMR (CDCl₃): δ 7.34-7.07 (m,18H), 4.68 (m,6H), 3.57 (m,4H), 3.17 (m,2H), 2.96 (m,2H), 2.79 (m,2H).
- 10 Example 13T: MS: 617(M+H,100%); NMR (CDCl₃,300MHz): δ 7.3(m,10H), 6.7(m,4H), 6.35(m,2H), 4.75(d,2H), 3.5(s,4H), 3.3(m,4H), 3.0(m,6H), 2.7(m,4H), 1.9(m,4H).
- 15 Example 13U: m.p. 264-266 °C; MS: 609(M+H,100%); NMR (CDCl₃,300MHz): δ 8.6(d,2H), 7.95(d,2H), 7.8(d,2H), 7.6(d,2H), 7.3(m,14H), 5.1(d,2H), 3.7(m,4H), 3.2(m,6H).
- 20 Example 13V: MS: 521(M+H,100%); NMR (CDCl₃,300MHz): δ 8.8(s,2H), 7.2(m,12H), 4.15(s,2H), 3.8(t,2H), 3.6(m,4H), 2.9(m,4H).
- Example 13W: m.p. 182-184 °C; MS: 687(M+H,100%);
- 25 NMR (CDCl₃,300MHz): δ 7.2(m,10H), 6.95(d,2H), 6.2(d,2H), 4.9(d,2H), 4.15(d,2H), 3.9(bs,2H), 3.7(d,2H), 3.0(m,6H), 1.45(s,18H).
- Example 13X: MS: 547(M+H,100%); NMR
- 30 (CDCl₃,300MHz): δ 7.2(m,10H), 6.2(d,2H), 6.0(d,2H), 4.8(d,2H), 4.5(bs,2H), 3.8(s,2H), 3.5(d,2H), 3.1(m,6H).
- Example 13Y: MS: 507 (M+1, 100%); NMR
- 35 (CDCl₃,300MHz): δ 7.0-7.4 (m, 20H), 4.7-4.8 (d,

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2H), 4.5 (m, 2H), 3.45 (m, 4H), 3.1 (m, 4H), 2.9 (m, 2H).

5 Example 13Z: MS: 577((M+H)⁺, 100%); 594((M + NH₄)⁺, 80%). NMR (CDCl₃): δ 8.08(s 1H), 7.48(dd, 1H, J = 8.4, 1.9 Hz), 7.25-7.38(m, 3H); 7.19(d, 1H, J = 8.4 Hz), 7.00-7.03(m, 2H), 3.96(ABq, 2H), 3.73(d, 1H, J = 1.1 Hz), 3.50(dd, 1H, J = 11.4, 1.1Hz), 3.23(br.s., 1H), 2.98(ABx, 2H).

10 Example 14A: NMR (CDCl₃): δ 8.27(dd, 1H, J = 4.9, 2.0 Hz), 7.66(dd, 1H, J = 7.5, 2.0 Hz), 7.14-7.27(m, 4H), 6.68(dd, 2H, J = 7.5, 3.6 Hz), 4.16(d, 1H, J = 0.8 Hz), 3.90(ABq, 2H), 3.72(dd, 1H, J = 11.3, 1.1Hz), 3.14(br.s., 1H), 2.87(ABx, 2H).

20 Example 14B: MS: 685((M+H)⁺, 100%). NMR (CDCl₃): δ 8.34(dd, 1H, J = 4.8, 1.9 Hz), 7.16-7.37(m, 4H), 6.93(dd, 1H, J = 7.6, 4.8 Hz), 6.87(dd, 2H, J = 7.5, 1.7 Hz), 4.14(s, 1H), 3.90(ABq, 2H), 3.68(td, 1H, J = 6.3, 0.7 Hz), 2.99(d, 2H, J = 6.3 Hz), 2.43(br.s., 1H).

25 Example 14C: MS: 541((M+H)⁺, 100%). NMR (CD₃OD): δ 7.53(dd, 1H, J = 9.5, 2.6 Hz), 7.15-7.26(m, 3H), 7.06(d, 1H, J = 2.6 Hz), 6.96-6.99(m, 2H), 6.42(d, 1H, J = 9.5 Hz), 3.80(s, 1H), 3.71(ABq, 2H), 3.62(dd, 1H, J = 11.7, 0.7 Hz), 2.92(ABx, 2H).

Example 14D: MS: 458(M+H⁺, 100%).

Example 14E: MS: 543(M+H⁺, 100%) NMR (CDCl₃): δ
7.4-7.1(m, 5H), 4.42(d, J = 17.5 Hz, 1H), 4.10(s,
5 1H), 3.83(d, J = 10.6 Hz), 3.2-3.0(m, 2H), 2.82(d,
J = 17.6 Hz, 1H), 2.36(s, 1H), 2.14(m, 2H), 1.5-
1.2(m, 6H), 0.85(t, J = 7.0 Hz, 3H).

Example 14F: MS: 739.5(M+H⁺) NMR (CDCl₃): δ
10 7.4-7.0(m, 15H), 3.90(s, 1H), 3.3-3.1(m, 2H),
3.04(m, 2H), 2.43(dd, J = 14.7, 5.7 Hz, 1H),
2.17(s, 1H), 1.824(m, 1H), 1.26(t, J = 6.0 Hz, 1H),
1.02(dd, J = 9.0, 5.1 Hz, 1H)

15 Example 14G: MS: 479(M+H⁺, 100%) NMR (CDCl₃): δ
7.24(bs, 5H), 7.15-6.9(m, 3H), 6.72(d, J = 7.5 Hz,
2H), 4.15(s, 1H), 4.06(d, J = 10.5 Hz, 1H), 3.5-
3.2(m, 2H), 2.98(s, 1H).

20 Example 14H: MS: 587(M+H⁺, 100%) NMR (CDCl₃): δ
7.4-6.9(m, 10H), 3.92(s, 1H), 3.8-3.6(m, 2H),
3.03(m, 2H), 2.50(bs, 1H), 2.19(dd, J = 14.3, 7.0
Hz, 1H), 1.57(m, 1H), 1.0-0.7(m, 2H).

25 Example 14I: MS: 457(M+H⁺, 100%), 474(M + NH₄⁺,
15%) NMR (CDCl₃): δ 7.4-6.9(m, 15H), 4.69(d, J =
6.2 Hz, 1H), 3.95(m, 2H), 3.77(m, 2H), 3.37(m, 1H),
3.25-3.0(m, 3H), 2.15(b, 1H), 1.61(m, 2H), 1.24(m,
1H), 1.0-0.7(m, 2H).

30 Example 14J: NMR (CDCl₃): δ 7.42(d, 2H, J = 7.3
Hz), 7.22-7.38(m, 6H), 7.11(d, 2H, J = 6.6 Hz),
6.54(s, 1H), 6.35(s, 1H), 5.31(s, 2H), 3.84(ABq,
2H), 3.72(s, 1H), 3.49(d, 1H, J = 10.4 Hz), 2.90-
35 3.01(m, 2H), 2.39(s, 3H).

Example 14K: NMR ((CD₃)₂SO): δ 7.00-7.33(m, 7H),
3.61(s, 1H), 3.55(ABq, 2H), 3.45(d, 1H, J = 11.0
Hz), 2.95(ABx, 2H), 2.09(s, 3H).

5

The structures of the Examples below are shown
in Table 2e.

10

Ketal formation: Preparation of Triacetone
(XXVIa):

Lithium borohydride (1.2 gr, 56.2 mmol) was
added in four portions to a suspension of L-
mannonic-g-lactone (5 gr, 28.1 mmol) in methanol
15 (250 mL) at 0 °C over 10 min. Ice bath was removed
and reaction stirred at room temperature for 30 min.
Reaction was quenched at 0 °C with 2N HCl. Solvent
was evaporated and residue taken up in acetone (75
mL) to which 2,2-dimethoxypropane (20 mL, 168.6
20 mmol) and camphorsulphonic acid (20 gr, 84.3 mmol)
were added in four portions. Reaction becomes clear
for a few minutes and then a precipitate forms.
Reaction stirred at room temperature for 14 h.
Solvent volume then reduced by 2/3 at reduced
25 pressure and then poured into EtOAc, washed with
saturated NaHCO₃, dried (MgSO₄) and concentrated.
Solid residue taken up in hexane and filtered thru a
pad of silica gel. Filtrate concentrated to give
triacetone (XXVIa) as a yellowish solid (7.1 gr,
30 80%). m.p. 72-74 °C; MS: 303 (M+H, 100%); NMR
(CDCl₃, 300MHz): δ 4.25 (m, 2H), 4.15 (m, 2H), 4.05
(m, 4H), 1.5 (s, 6H), 1.45 (s, 6H), 1.4 (s, 6H).

35

Selective Acetone Deprotection: Preparation of
Tetraol (XXVIb):

Compound (XXVIA) (14 gr) in 70% AcOH (200 mL) was stirred at 45 °C for 2 h. Solvent removed at reduced pressure with a bath temperature of 45 °C. Residue recrystallized from ether. Mother liquors concentrated and chromatographed (silica, 10% methanol in methylene chloride) to give the desired product as a white solid (8.2 gr, 80%). m.p. 91-93 °C; $[\alpha]_D = -26.40$ (c=3, H₂O); MS: 240 (M+NH₄, 100%); NMR (CDCl₃, 300MHz): δ 3.95 (m, 6H), 3.75 (m, 4H), 2.5 (bs, 2H), 1.4 (s, 6H).

Epoxide Formation: Preparation of Diepoxide (XXVIC):

A solution of Compound (XXVIB) (1 g, 4.5 mmol) in pyridine (5 mL) was cooled to -20 °C and treated with p-toluenesulfonyl chloride (1.89 g, 10 mmol). Stirring continued at -20 °C for 20 min, 0 °C for 20 min, and 23 °C for 20 min. The reaction was then diluted with methylene chloride and washed with 2N HCl and NaHCO₃. The organic extract was dried over MgSO₄ and concentrated. The crude product was then taken up in methanol (14 mL) and cooled to 0 °C. Next, K₂CO₃ (3.11 g, 22 mmol) was added and the reaction stirred at room temperature for 30 min. The methanol was then stripped off (do not evaporate to dryness, epoxide is volatile) and the crude was washed with water, extracted with ether, dried over MgSO₄, filtered, and concentrated. The compound was purified on SiO₂ and eluted with 30-60% ether/petroleum ether to afford the diepoxide (0.63 g, 75%) as an oil. NMR (CDCl₃, 300MHz): δ 3.81 (m, 2H), 3.10 (m, 2H), 2.80 (t, 2H), 2.68 (m, 2H), 1.40 (s, 6H, CH₃).

Opening of Epoxide: Prepartation of Diol (XXVID):

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To a suspension of cuprous bromide-dimethyl sulfide complex (1.8 g, 8.7 mmol) in anhydrous THF (5 mL) at -20 °C was added 8.5 ml benzylmagnesium chloride (2M in THF, 17 mmol). Reaction stirred at -20 °C for 30 min and at 0 °C for 1 h. Next, Compound (XXVIc) (0.54 g, 3 mmol) in THF (5 mL) was added, and the reaction stirred at 0 °C for 1 h. The excess reagent was quenched with saturated NH₄Cl solution and allowed to warm to room temperature. The contents were then washed with water and brine, extracted with ether, dried over MgSO₄, filtered, and concentrated. Crude material was then purified by flash chromatography (30-60% ether/petroleum ether) to yield 0.84 g (78%) of an oil. MS: 371 (M+H, 66%); NMR (CDCl₃, 300MHz): δ 7.2-7.4 (m, 10H), 4.65 (s, 2H), 3.6-3.8 (m, 4H), 2.6-3.0 (m, 4H), 1.8-2.2 (m, 4H), 1.4 (s, 6H).

Hydroxyl Displacement: Preparation of Diazide (XXVIe):

To a solution of Compound (XXVIId) (0.48 g, 1.3 mmol) and triphenyl phosphine (1.0 g, 3.9 mmol) in THF (5 mL) at 0 °C was added diethylazodicarboxylate (0.61 mL, 3.9 mmol) and diphenylphosphorylazide (0.84 mL, 3.9 mmol). Contents were allowed to warm to room temperature in the ice bath for 1 h. The excess reagents were quenched by the addition of methanol (0.2 mL, 5 mmol) at 0 °C. The mixture was then stirred at room temperature for 30 min and then concentrated to a small volume. Crude was then purified on SiO₂ using 1:40 ethyl acetate/hexane giving 0.245 g (45%) of an oil. MS: 438 (M+NH₄, 8%); NMR (CDCl₃, 300MHz): δ 7.2-7.4 (m, 10H), 4.18 (s, 2H), 2.7-3.0 (m, 6H), 2.0-2.3 (m, 4H), 1.58 (s, 6H).

Reduction of Diazide (XXVIe):

To Compound (XXVIe) (0.245 g, 0.58 mmol) in ethanol (6 mL) under N₂ was added 10% Pd/C (73.5 mg, 30%/weight). Reaction stirred under H₂ atmosphere at room temperature overnight. Crude was then filtered through celite and concentrated. 0.21 g (98%) of the diamine was collected as an oil and taken onto next step without further purification. MS: 369 (M+H, 100%); NMR (CDCl₃, 300MHz): δ 7.05-7.3 (m, 10H), 3.9 (bs, 2H), 3.05 (bs, 4H), 2.8 (m, 2H), 2.6 (m, 4H), 1.7 (bs, 4H), 1.35 (s, 6H).

Cyclization of the Diamine: Formation of Cyclic Urea (XXVI f):

The diamine (0.21 g, 0.57 mmol) was dissolved in methylene chloride (50 mL) and carbonyldiimidazole (0.102 g, 0.63 mmol) was added and the reaction stirred at 23 °C overnight. The solution was then concentrated and purified on SiO₂ using 75% ethyl acetate/hexane as eluent which gave 85mg (38%) of (XXVI f) as a foam. MS: 395 (M+H, 100%); NMR (CDCl₃, 300MHz): δ 7.0-7.2 (m, 10H), 3.6-4.0 (m, 4H), 3.6-2.7 (m, 4H), 1.8-1.9 (m, 4H), 1.3 (s, 6H).

Alkylation of the Cyclic Urea (XXVI f):

To Compound (XXVI f) (85mg, 0.22 mmol) in dry DMF (3 mL) was added 60% NaH (0.07 g, 1.7 mmol). The solution was stirred for 5 min at room temperature. Next, benzyl bromide (0.1 mL, 0.86 mmol) was added and the reaction stirred at 23 °C overnight. Reaction was then quenched with methanol (several drops), washed with H₂O, extracted with ether, dried (MgSO₄), and concentrated. Crude was then purified on silica gel using 1:1 hexane/ethyl acetate affording 0.03 g (25%) of the bis-alkylated urea as a foam. MS: 575 (M+H, 100%); NMR (CDCl₃, 300MHz): δ

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7.1-7.4 (m, 20H), 5.1 (d, 2H), 4.0 (d, 2H), 3.75 (bs, 2H), 3.6 (m, 2H), 2.7 (m, 2H), 2.6 (m, 2H), 1.9-2.0 (m, 4H), 1.25 (s, 6H).

Deprotection of Acetonide: Preparation of Example

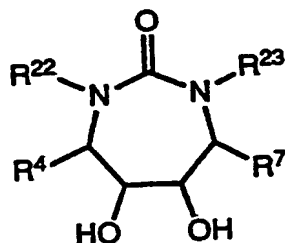
16A:

5

To above prepared bis-alkylated cyclic urea (0.03 g, 0.05 mmol) in THF (2 mL) at room temperature was added several drops of concentrated HCl. Reaction stirred at room temperature for 2 h. Reaction was then washed with 1 N NaOH, extracted with ethyl acetate, dried (MgSO₄), and concentrated. Chromatography (silica, 1-5% methanol in methylenechloride) gave 0.024 g (85%) of example 16A as a foam. MS: 535 (M+H, 100%); NMR (CDCl₃, 300MHz): δ 7.1-7.3 (m, 20H), 5.15 (d, 2H), 3.9 (d, 2H), 3.5 (bs, 2H), 3.3-3.4 (m, 2H), 2.7-2.8 (m, 2H), 2.5-2.6 (m, 2H), 2.0-2.1 (m, 6H).

20

Table 2e



Ex. No.	Stereo	R ²²	R ²³	R ⁴ = R ⁷	HPLC K _i	Note
	<u>2:3:4:5</u>					
16A	RSSR	benzyl	benzyl	phenethyl	++	1
16B	RSSR	allyl	allyl	phenethyl	+	1
16C	RSSR	allyl	allyl	isopropyl	++	2
16D	RSSR	cyclopropyl methyl	cyclopropyl methyl	isopropyl	++	2

16E	RSSR	allyl	allyl	methyl	+	2
16F	RSSR	allyl	n-butyl	methyl	+	2
16G	RSSR	benzyl	benzyl	methyl	+	2
16H	RSSR	2-naphthyl- methyl	2-naphthyl- methyl	methyl	+	2
16I	RSSR	allyl	allyl	isobutyl	+++	2
16J	RSSR	cyclopropyl methyl	cyclopropyl methyl	isobutyl	+++	2

Note 1: Prepared as in Scheme 4.

Note 2: Prepared as compounds in Table 1A.

- 5 Listed below are physical data for
representative compounds of the invention.

Example 16A: MS: 535 (M+1, 100%); NMR
(CDCl₃, 300MHz): δ 7.1-7.3 (m, 20H), 5.15 (d, 2H),
10 3.9 (d, 2H), 3.5 (bs, 2H), 3.3-3.4 (m, 2H), 2.7-2.8
(m, 2H), 2.5-2.6 (m, 2H), 2.0-2.1 (m, 6H).

Example 16B: MS: 435 (M+1, 100%); NMR
(CDCl₃, 300MHz): δ 7.1-7.3 (m, 10H), 5.8 (m, 2H),
15 5.15 (s, 2H), 5.1 (d, 2H), 4.5-4.6 (m, 2H), 3.8 (s,
2H), 3.3-3.5 (m, 4H), 2.5-2.9 (m, 4H), 2.2 (m, 2H),
2.0 (m, 4H).

Example 16C: MS: 312 (22, M + 2), 311 (100, M +
20 1), 267 (1). HRMS: Calc. 311.2334. Found: 311.2330.
NMR (CDCl₃): δ 5.81(m, 2H), 5.2(m, 4H), 4.4(m, 2H),
4.0(br s, 2H), 3.4 (m, 2H), 3.1(br s, 2H), 3.0(m,
2H), 2.4(m, 2H), 1.2(d, 6H), 0.9(d, 6H).

25 Example 16D: MS: 341(4), 340 (25), 339 (100, M +
1), 321 (1), 295 (2), 256 (2). HRMS: Calc.
339.2647. Found: 339.2652. NMR (CDCl₃): δ 4.2(brs,
2H), 3.65(m, 2H), 3.20(m, 4H), 2.6(m, 4H), 1.2(d,
6H), 1.0(m, 2H), 0.9(d, 6H), 0.5(m, 4H), 0.2(m, 4H).

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Example 16E: MS: 256 (15, M + 2), 255 (100, M + 1). HRMS: Calc. 255.1706. Found: 255.1708. NMR (CDCl₃) δ 5.8(m, 2H), 5.2(m, 4H), 4.0(m, 2H), 3.8(br s, 2H), 3.65(m, 2H), 3.4(m, 2H), 2.8(m, 2H), 1.2(d, 6H).

Example 16F: MS: 272 (16, M + 2), 271 (100, M + 1). HRMS: Calc. 271.2021. Found: 271.2036. NMR (CDCl₃): δ 5.8(m, 1H), 5.2(m, 2H), 4.0(m, 2H), 3.8(br s, 2H), 3.6-3.3(3H), 3.0(m, 2H), 2.45(m, 2H), 1.35(m, 2H), 1.2(d, 6H), 0.9(m, 4H).

Example 16G: MS: 356 (23, M + 2), 355 (100, M + 1). HRMS: Calc. 355.2021. Found: 355.2012. NMR (CDCl₃): δ 7.4-7.2(m, 10H), 4.9(d, 2H), 4.2(d, 2H), 3.6(br s, 2H), 3.3(m, 2H), 2.2(m, 2H), 1.2(d, 6H).

Example 16H: mp 236-238°C. MS: 456 (29, M + 2), 455 (100, M + 1), 315 (10), 158 (5). HRMS: 455.2334. Found: 455.2333. NMR (CDCl₃): δ 7.9-7.8(m, 8H), 7.6-7.45(m, 6H), 5.0(d, 2H), 4.4(m, 2H), 3.6(br s, 2H), 3.4(m, 2H), 1.9(m, 2H), 1.2(m, 2H).

Example 16I: MS 339 (100, M+1), 311 (4). HRMS 339.265377 (calc. mass = 339.264768). NMR (CDCl₃): δ 5.91-5.79 (m), 5.29-5.18 (m, 4H), 4.53-4.46 (m, 2H), 3.76 (s, 2H), 3.41-3.33 (m, 4H), 2.13 (s, 2H), 1.81-1.71 (m, 2H), 1.71-1.62 (m, 2H), 1.45-1.35 (m, 2H), 0.90 (t, 12H).

Example 16J: m.p. 137-139 °C. MS 367 (100, M+1). HRMS 367.295357 (calc. mass = 367.296068). NMR (CDCl₃): δ 3.94 (s, 2H), 3.82 (d, 2H), 3.77 (d, 2H), 3.48 (d, 2H), 2.91 (bs, 2H), 2.63 (d, 2H), 2.61 (d, 2H), 1.99 (m, 2H), 1.70-1.61 (m, 2H), 1.43 (m, 2H).

1.26 (s, 2H), 1.07-0.98 (m, 2H), 0.91 (t, 12H), 0.52 (m, 2H), 0.22 (m, 2H).

5 The structures of the Examples below are shown in Table 2f-h.

Synthesis of Thiourea (XXVIIa):

10 Diaminodimem Compound (XXIb) (22.45g, 47.1 mmol) was dissolved in 200 mL of tetrahydrofuran and to this solution was added 9.23g (51.8 mmol) of thiocarbonyl diimidazole. After stirring the mixture for 18 hours at room temperature TLC (10:1:10 ethyl acetate: ethanol: 15 hexane) indicated complete reaction. The reaction mixture was taken to dryness and the solid residue purified by flash chromatography (silica gel, 250g, 1:1 ethyl acetate: hexane) to provide solid which was trituated with hexane to provide 17.8g (73% yield) of 20 XXVIIa as a white solid.

Synthesis of Compound (XXVIIb):

25 Compound (XXVIIa) (3.108g, 6mmol) was dissolved in 15ml acetonitrile and to this solution was added methyl iodide 1.5ml (24mmol) via syringe and stirred at room temperature for one hour. The contents were then taken to dryness. The residue was dissolved in 30ml dimethylformamide and to this solution, cooled in a 0°C 30 ice bath, was added NaH (60% in oil) 720mg (18mmol) slowly (EVOLUTION!). The contents were stirred at room temperature for 30 minutes. The mixture was cooled in a 0°C ice bath and benzyl bromide (2.052g, 12mmol) was added via syringe and stirred at room temperature for 18 35 hours. TLC (2:3 EtOAc:Hexane R_f =0.25) indicated a complete reaction. The reaction was worked up by

- diluting with water (300ml) and extracting with diethyl ether (3x50ml). The organic layer was dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel (200g; 2:3 EtOAc:Hexane) to provide 2.923g (78.2% yield) of XXVIIb as a colorless oil.

Synthesis of Compounds (XXVIIc) and (XXVIIId):

- Compound (XXVIIb) (2.900g, 4.65mmol) was dissolved in 25ml pyridine and to this solution was added 742mg (4.65mmol) benzylhydroxylamine hydrochloride. The contents were refluxed in a 125°C oil bath for 18 hours. (Caution: Methyl mercaptan is a by-product and the reaction should be vented to a Clorox scrubber). TLC indicated a complete reaction. The reaction was diluted with 150ml dichloromethane. The organic layer was washed with 1N HCl (2x300ml) followed by sat. sodium bicarbonate solution (100ml). It was separated and dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel (130g; using 1:3 EtOAc:Hexane) to provide 584mg (18.0% yield) of Compound (XXVIIc) as a colorless oil. 1:2 EtOAc:Hexane was used to provide 2.113g of a side product thiourea (XXVIIId).

Synthesis of Oxime (XXVIIe):

- Compound (XXVIIc) (584mg, 0.84mmol) was dissolved in 5ml dimethylformamide and to this solution, cooled in a 0°C ice bath, was added NaH (60% in oil) 80mg (2mmol) slowly (EVOLUTION!). The contents were stirred at room temperature for 30 minutes. The mixture was cooled in a 0°C ice bath and benzyl bromide (0.24ml, 2mmol) was added via syringe and stirred at room temperature for 18 hours. TLC (1:3 EtOAc:Hexane R_f=0.26) indicated a

complete reaction. The reaction was worked up by diluting with water (50ml) and extracting with diethyl ether (2x25ml). The organic layer was dried over magnesium sulfate and the filtrate taken to dryness.

- 5 The residue was purified on SiO₂ gel (33g; 1:3 EtOAc:Hexane) to provide 491mg (74.2% yield) of a colorless oil.

Example 18A

10

Compound (XXVIId) (450mg, 0.57mmol) was placed in a 25ml R.B. Flask and cooled in a 0°C ice bath. To this flask was added 4M HCl in dioxane (5ml, 20mmol) and the mixture stirred at room temperature for 18 hours. TLC

- 15 (2:3 EtOAc:Hexane R_f=0.29) indicated a complete reaction. The mixture was worked up by quenching in sat. sodium bicarbonate solution (50ml) and extracting with dichloromethane (2x50ml). The organic extracts were dried over magnesium sulfate and the filtrate taken
20 to dryness. The residue was purified on SiO₂ gel (33g; 2:3 EtOAc:Hexane) to provide 246mg (70.5% yield) of Example 18A as a waxy solid.

Example 17A

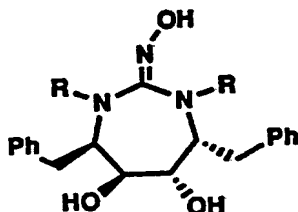
25

Example 18A (160mg, 0.26mmol) was dissolved in 5ml ethanol. To this mixture was added 50mg of 10% palladium hydroxide on Carbon and the suspension stirred for 18 hours under hydrogen (1
30 atm). TLC (10:1:10 EtOAc:EtOH: Hexane R_f=0.3)

- indicated a complete reaction. The suspension was filtered through a celite pad and the filtrate taken to dryness. The residue was purified on SiO₂ gel (33g; 10:1:10 EtOAc:EtOH:Hexane) to provide 97mg of
35 Example 17A (69.5% yield) as a white solid.

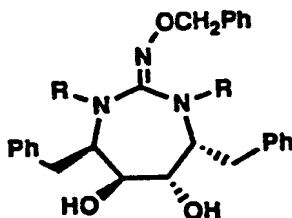
Example 19A

Compound (XXVIId) (500mg, 0.82mmol) was placed in a 25ml R.B. Flask and cooled in a 0°C ice bath. To this flask was added 4M HCl in dioxane (7.5ml, 30mmol) and the mixture stirred at room temperature for 18 hours. TLC (1:2 EtOAc:Hexane $R_F=0.29$) indicated a complete reaction. The mixture was worked up by quenching in sat. sodium bicarbonate solution (50ml) and extracting with dichloromethane (2x50ml). The organic extracts were dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel (33g; 1:2 EtOAc:Hexane) to provide 181mg (51.1% yield) of Example 19A as a white solid.

Table 2f

<u>Ex. No.</u>	<u>R</u>	<u>HPLC</u> <u>Ki</u>	<u>IC₅₀</u>	<u>m.p.</u> °C	<u>MS</u> <u>M+H</u>
17A	CH ₂ C ₆ H ₅	++		120.1	522.275

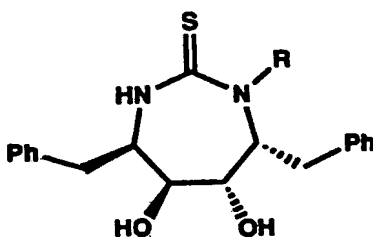
Table 2g



Ex. No.	R	HPLC Ki	IC ₉₀	m.p. °C	MS M+H
18A	CH ₂ C ₆ H ₅	+		59.5	612.322

5

Table 2h



10

Ex. No.	R	HPLC Ki	IC ₉₀	m.p. °C	MS M+H
19A	CH ₂ Ph	+++	+++	74.0	

The structures of the Examples below are shown in Table 2i.

15

Acetylation of Diol: Compound (XXVIIIa):

Example 1X (3.517g, 7.58mmol) was dissolved in 25ml pyridine and to this solution, cooled in a 0°C ice bath, was added 350mg 4-Dimethylaminopyridine and 7.16ml (75.85mmol) acetic anhydride. The contents were stirred at room temperature for 18

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hours. TLC (1:4 EtOAc:Hexane $R_f=0.3$) indicated a complete reaction. The reaction was diluted with 250ml dichloromethane. The organic layer was washed with 1N HCl (2x300ml) followed by sat. sodium bicarbonate solution (100ml). It was separated and dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel (200g; 1:5 EtOAc:Hexane) to provide 2.632g (67.0%) of XXVIIIa as a white solid.

10 Nitration of Benzyl Group: Compound (XXVIIIb) and (XXVIIIc):

Compound (XXVIIIa) (518mg, 1mmol) was dissolved in 4ml acetonitrile and to this solution, cooled in a -40°C dry ice-acetone bath, was added 4.4ml (2.2mmol) 0.5M Nitronium tetrafluoroborate in sulfolane. The contents were stored in a -40°C freezer for 18 hours. TLC indicated a complete reaction. The reaction was diluted with 100ml ether and washed with water (2x50ml). The organic layer was dried over magnesium sulfate and the filtrate taken to dryness. The residue was purified on SiO₂ gel (75g; 1:3 EtOAc:Hexane for XXVIIIb, 1:2 EtOAc:Hexane for XXVIIIc) to provide 106mg (17.4% yield) of XXVIIIb as a white solid and 159mg (26.2% yield) of XXVIIIc as a white solid.

Example 20A

30 Compound (XXVIIIb) (106mg, 0.174mmol) was dissolved in 5ml methanol and to this solution was added 0.5ml 0.5M sodium methoxide in methanol via syringe. The contents were stirred at room temperature for 30 minutes. TLC indicated a complete reaction. The mixture was quenched by

adding 500mg of AG50W-X8 acid resin and stirring the suspension at room temperature for 5 minutes. The filtrate was taken to dryness and the residue purified on SiO₂ gel (33g; 1:2 EtOAc:Hexane) to provide 43mg (47.1% yield) of Example 20A as a white solid.

Example 20B

Compound (XXVIIIc) (159mg, 0.261mmol) was dissolved in 5ml methanol and to this solution was added 0.5ml 0.5M sodium methoxide in methanol via syringe. The contents were stirred at room temperature for 30 minutes. A white precipitate started forming after 15 minutes. TLC indicated a complete reaction. The mixture was quenched by adding 500mg of AG50W-X8 acid resin and stirring the suspension at room temperature for 5 minutes. 10ml dichloromethane was then added to solubilize the solid. The filtrate was taken to dryness and the residue provided 111mg (81.1% yield) of Example 20B as a white solid.

Example 20E

Example 20A (100mg, 0.191mmol) was dissolved in 5ml ethanol. To this mixture was added 50mg of 5% palladium on Carbon and the suspension stirred for 18 hours under hydrogen (1 atm). TLC indicated a complete reaction. The suspension was filtered through a celite pad and the filtrate taken to dryness. The residue provided 47mg (53.0% yield) of Example 20E as a white solid.

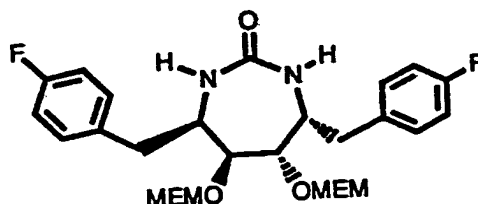
Example 20F

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Example 20B (100mg, 0.191mmol) was suspended in 5ml ethanol. To this mixture was added 50mg of 5% palladium on Carbon and the suspension stirred for 18 hours under hydrogen(1 atm). The starting material went into solution as the reaction progressed. TLC indicated a complete reaction. The suspension was filtered through a celite pad and the filtrate taken to dryness. The residue provided 49mg (55.2% yield) of example 20F as a white solid.

Example 20G

A. Synthesis of 4-Fluorobenzyl Cyclic Urea (XXXI):



(XXXI)

The synthesis of 4-fluorobenzyl cyclic urea is outlined in Scheme 7. N-acetyl-D-4-fluorophenylalanine methyl ester (23.9 g, 0.1 mol), obtained using the procedure of M.J. Burk (*J. Am. Chem. Soc.* **1991**, *113*, 8518), was dissolved in 40 mL of acetic acid and treated with 100 mL of concentrated HCl, 40 mL of water and heated to reflux for 5 hrs. The solution was cooled to room temperature and then made basic (pH = 10) with 50% NaOH while cooling in an ice bath. Benzyl chloroformate (25 mL, 29 g, 0.17 mol) and NaOH are added in four portions and the solution is

maintained alkaline by the addition of NaOH. The mixture is then stirred at rt for 30 min. The alkaline solution is extracted with ether (2 X 500 mL) and the solution acidified with conc HCl to pH

- 5 1. The precipitate is extracted into methylene chloride and dried over MgSO_4 . The solution is filtered and concentrated to give 20 g of the N-Cbz-D-4-fluorophenylalanine as a white solid that is used without further purification.

- 10 A solution of N,O-dimethylhydroxylamine hydrochloride (8.0 g, 0.082 mol) in DMF is prepared by gentle warming. The solution is allowed to cool slightly and treated with N-methylmorpholine (8.2 g, 0.082 mol) and diluted with THF to facilitate
15 transferring of the resulting thick suspension.

- A solution of N-Cbz-D-4-fluorophenylalanine (20 g, 0.063 mol) in THF is treated with N-methylmorpholine (9.0 g 0.09 mol) and cooled to 0°C in an ice bath. To the stirred cold solution is
20 added isobutyl chloroformate (8.6 g, 0.063 mol) in small portions over a period of 10 mins. Then the solution of N,O-dimethylhydroxylamine in DMF prepared above is added and the reaction mixture is stirred for 20 mins. Most of the solvent is removed
25 on a rotary evaporator and the residue is partitioned between water and methylene chloride. The organic layer is washed successively with 1 N HCl, 1 N NaOH, water, brine and then dried over MgSO_4 . The solution is then filtered and
30 concentrated and the residue chromatographed on silica gel (50% EtOAc/Hex) to give 16 g of the amide.

- Using the procedure of by J-A. Fehrentz and B. Castro (*Synthesis*, 1983, 676) 11 g (0.031 mol) of
35 N-Cbz-D-4-fluorophenyl-alanine N,O-dimethylhydroxylamide was converted to 9.0 g of N-

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Cbz-D-4-fluorophenylalaninal obtained as a thick oil that was used without further purification.

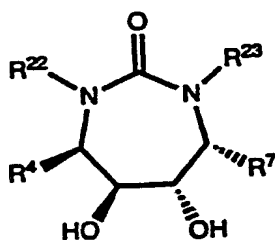
N-Cbz-D-4-fluorophenylalaninal (9.0 g, 0.031 mol) was converted, using procedure 1, to
 5 (2R,3S,4S,5R)-2.5-bis(N-Cbz-amino)-3,4-dihydroxy-1,6-di(4-fluorophenyl)hexane (4 g) obtained as a white solid. MS: (CI, NH₃) (M+H)⁺ = 605.

The (2R,3S,4S,5R)-2.5-bis(N-Cbz-amino)-3,4-dihydroxy-1,6-di(4-fluorophenyl)hexane (4.0 g,
 10 0.0066 mol) was converted, as described in procedure 4, to 1.3 g of the 4-fluorobenzyl cyclic urea (XXXI) obtained as a white solid. MS: (CI, NH₃) (M+H)⁺ = 539.3

15 B. The 4-fluorobenzyl cyclic urea (XXXI) (270 mg, 0.5 mmol) was alkylated with 3-benzoyloxybenzyl chloride (350 mg, 1.5 mmol) according to general procedure 5. The resulting intermediate was
 20 dissolved in THF and hydrogenated for 12 hours (200 mg 10% Pd/C, 55 psi) to remove the benzyl protecting groups. The MEM group was then removed, according to general procedure 5, to give, after chromatography on HPLC (silica gel, 10% MeOH/CHCl₃), 140 mg of Example 20G as a white foam.
 25 MS: (CI, NH₃) (M+H)⁺ = 575.2 (100%).

Table 2i

30



Ex. No.	R ²² , R ²³	R ⁴	R ⁷	HPLC Ki	IC ₅₀	m.p. °C	MS M+H
20A	cyclopropyl-methyl	4-nitro-benzyl	2-nitro-benzyl	+++	+++	209.8	525.234
20B	cyclopropyl-methyl	4-nitro-benzyl	4-nitro-benzyl	+++	+++	227.5	525.234
20C	n-butyl	4-nitro-benzyl	2-nitro-benzyl	+++	+++	165.0	529.266
20D	n-butyl	4-nitro-benzyl	4-nitro-benzyl			245.0 (dec)	529.266
20E	cyclopropyl-methyl	4-amino-benzyl	2-amino-benzyl	+++	+++		465.286
20F	cyclopropyl-methyl	4-amino-benzyl	4-amino-benzyl	++	+++		465.286
20G	3-hydroxy benzyl	4-fluoro benzyl	4-fluoro benzyl	+++			575.2
20H	cyclopropyl-methyl	4-fluoro benzyl	4-fluoro benzyl	+++			471.2
20I	4-hydroxy methylbenzyl	4-fluoro benzyl	4-fluoro benzyl	+++			603.2
20J	4-acetyl benzyl	benzyl	benzyl	++			608 (M+NH ₄)
20K	4-fluoro methylbenzyl	benzyl	benzyl	+++	+++		571

The structures of the Examples below are shown
 5 in Table 2j.

Example 21A

A. Synthesis of Aziridine Urea (XXXIIa):

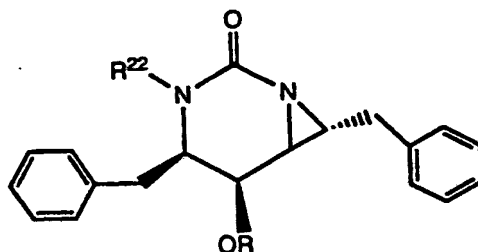
10 A solution of Example 1A (5.3 g, 0.016 mol) in pyridine was treated with acetic anhydride (3.3 g, 0.033 mol) and stirred at room temperature for 3 hrs. 10 mL of MeOH was added and the mixture was
 15 evaporated to dryness. The residue was extracted into methylene chloride and washed sequentially with water, 1 N HCl, brine, and dried over MgSO₄. The solution was filtered, concentrated and the residue chromatographed on silica gel
 (5%MeOH/CHCl₃) to give 2.0 g of the corresponding
 20 monoacetate product as a white solid. The solid obtained was dissolved in methylene chloride and

cooled in an ice bath under nitrogen. To this was added DAST (0.875 g, 0.005 mol) via syringe and the solution stirred for 10 mins. The mixture was quenched with sat'd NaHCO_3 and the organic layer washed with water and brine. The solution is dried over MgSO_4 then filtered and concentrated to give 1.9 g of the acetate aziridine (XXXIIa) which is used without further purification.

5
10
15
B. The acetate aziridine (XXXIIa) (100 mg, 0.29 mmol) is dissolved in MeOH (2 mL) and treated with 1 N NaOH (0.5 mL) and stirred at rt for 30 min. The mixture is diluted with water (20 mL) and extracted into CH_2Cl_2 . The extract is washed with water and brine, dried over MgSO_4 then filtered and concentrated to give 30 mg of Example 21A as a white solid. MS: (CI, NH_3) $(\text{M}+\text{H})^+ = 309.0$

Example 21B and 21C

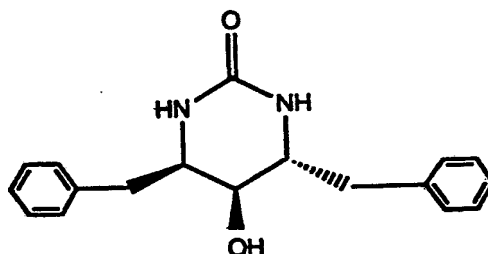
20 The acetate aziridine (XXXIIa) (200 mg, 0.57 mmol) is alkylated with benzyl bromide (120 mg, 0.69 mmol) according to Procedure 5 to give a mixture of products. This was HPLC chromatographed on silica gel (50% EtOAc/Hex) to give first 50 mg of Example 21B as a white solid. MS: (CI, NH_3) $(\text{M}+\text{H})^+ = 309.0$.
25 This was followed by 30 mg of Example 21C, obtained as a colorless oil. MS: (CI, NH_3) $(\text{M}+\text{H})^+ = 489.2$.



Ex No.	R ²²	R	K _i	IC ₉₀	MS M+H
21A	H	H	+	++	309.0
21B		Ac	+	++	441.0
21C			++		489.2
21D		H	+	++	399.1

The structures of the Examples below are shown in
5 Table 2k.

Preparation of the Cyclic Urea (XXXIIIIa):



10

A. Preparation of 4-Amino-2-(t-butoxycarbonylamino)-
1,5-diphenyl-3-(2-methoxyethoxymethyl)pentane.

15 A mixture of 595 mg (1.50 mmole) of 4-azido-2-(t-butoxycarbonylamino)-1,5-diphenyl-3-hydroxypentane (EP 0 402 646 A1), 10 ml of dioxane, 0.2 ml (1.75 mmole) of MEM chloride, and 0.32 ml (1.83 mmole) of

diisopropylethylamine was heated at 80°C for 16 hrs. Evaporated the solvent and purified the residue by flash chromatography on silica gel with 85:15 hexane-ethyl acetate to give 0.64 g (88%) of an oil. Mass spec
5 (M+H)⁺ = 485.2. This was reduced to the title compound with hydrogen using 100 mg of 10 % Pd on carbon in 60 ml of ethyl acetate and 0.6 ml of acetic acid in 49% yield.

10 B. Preparation of 2,4-diamino-1,5-diphenyl-3-hydroxypentane.

The product from Part A (218 mg) was dissolved in 2 ml of ice cold 1:1 trifluoroacetic acid -
15 dichloromethane. After 1 hr the solution was poured into a mixture of sodium bicarbonate solution and ethyl acetate. The ethyl acetate extract yielded 163 mg of the desired diamino compound.

20 C. Cyclization of the Diamine

The product from Part B (146 mg), 75 mg of carbonyl diimidazole, and 0.15 ml of diisopropylethylamine were dissolved in 2.5 ml of anhydrous THF and stirred at room
25 temperature for 16 hrs. The solvent was evaporated. The residue was purified by preparative TLC on silica gel with 90:10 dichloromethane - methanol to give 108 mg (69 %) of the cyclic urea. Mass spec (M+H)⁺ = 385.1.

30 N-Alkylation of the Cyclic Urea (XXXIIIa):

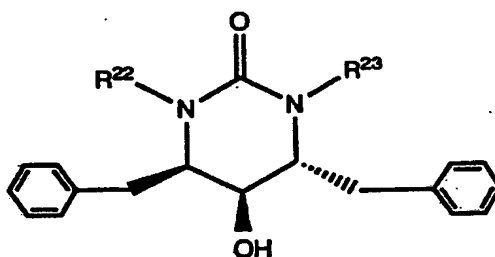
D. The product from Part C (93 mg) was dissolved in 2.5 ml of anhydrous DMF, and 100 mg of 60% NaH in mineral oil was added. The mixture was stirred for one
35 hr. m-Benzylloxbenzyl chloride (350 mg) was added, and the mixture was stirred for 16 hrs at room temperature.

Water and ethyl acetate were added. The ethyl acetate extract was washed with water, dried and evaporated. The residue was purified by prep TLC on silica gel with 60:40 hexane - ethyl acetate to give 105 mg (54%) of the desired bis-alkylated product. Mass spec (M+H)⁺ = 777.5

Deprotection of Protecting Groups (Example 22A):

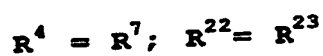
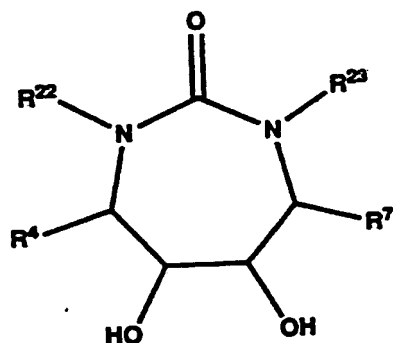
The product from part D (103 mg) was dissolved in 4N HCl/dioxane for 16 hrs. The solution was evaporated and purified by prep TLC on silica gel with 60:40 hexane - ethyl acetate. Mass spec (M+H)⁺ = 689.4. The purified material was hydrogenated for 16 hrs in the presence of 3 ml ethanol, 0.2 ml of acetic acid, and 35 mg of 10% Pd on carbon to give Example 22A. Mass spec (M+H)⁺ = 509.25; calculated, 509.24.

Table 2k



Ex No.	R ²²	R ²³	K _i	IC ₉₀	MS M+H
22A	m-(HO)- C ₆ H ₄ CH ₂ -	m-(HO)- C ₆ H ₄ CH ₂ -	++		509.25

Table 3



Example Number	$R^4 = R^7$	$R^{22} = R^{23}$
1	benzyl	allyl
2	1-pyrrolylmethyl	allyl
3	1-pyrazolylmethyl	allyl
4	1-imidazolylmethyl	allyl
5	1-indolylmethyl	allyl
6	1-triazolylmethyl	allyl
7	1-tetrazolylmethyl	allyl
8	2-pyridylmethyl	allyl
9	3-pyridylmethyl	allyl
10	4-pyridylmethyl	allyl
11	cyclohexylmethyl	allyl
12	2-naphthylmethyl	allyl
13	3-naphthylmethyl	allyl
14	2-thiophenylmethyl	allyl
15	4-(1-methyl)piperidinyl- methyl	allyl
16	"(3,4- methylenedioxyphenyl)methyl"	allyl
17	2-thienylmethyl	allyl
18	4-biphenylmethyl	allyl
19	pyrimidinylmethyl	allyl
20	2-benzothiazolylmethyl	allyl
21	2-benzothiophenylmethyl	allyl
22	2-thiomethylethyl	allyl
23	2-thiomethylmethyl	allyl
24	2-methylpropyl	allyl
25	2-methylbutyl	allyl

Table 3

26	3-methylbutyl	allyl
27	cyclopropylmethyl	allyl
28	cyclobutylmethyl	allyl
29	cyclopentylmethyl	allyl
30	p-hydroxyphenylmethyl	allyl
31	p-nitrophenylmethyl	allyl
32	p-aminophenylmethyl	allyl
33	"4-(N,N-dimethylamino)phenylmethyl"	allyl
34	benzyl	propyl
35	1-pyrollylmethyl	propyl
36	1-pyrazolylmethyl	propyl
37	1-imidazolylmethyl	propyl
38	1-indolylmethyl	propyl
39	1-triazolylmethyl	propyl
40	1-tetrazolylmethyl	propyl
41	2-pyridylmethyl	propyl
42	3-pyridylmethyl	propyl
43	4-pyridylmethyl	propyl
44	cyclohexylmethyl	propyl
45	2-naphthylmethyl	propyl
46	3-naphthylmethyl	propyl
47	2-thiophenylmethyl	propyl
48	4-(1-methyl)piperidinyl-methyl	propyl
49	"(3,4-methylenedioxyphenyl)methyl"	propyl
50	2-thienylmethyl	propyl
51	4-biphenylmethyl	propyl
52	pyrimidinylmethyl	propyl
53	2-benzothiazolylmethyl	propyl
54	2-benzothiophenylmethyl	propyl
55	2-thiomethylethyl	propyl
56	2-thiomethylmethyl	propyl
57	2-methylpropyl	propyl
58	2-methylbutyl	propyl
59	3-methylbutyl	propyl
60	cyclopropylmethyl	propyl
61	cyclobutylmethyl	propyl
62	cyclopentylmethyl	propyl
63	p-hydroxyphenylmethyl	propyl
64	p-nitrophenylmethyl	propyl
65	p-aminophenylmethyl	propyl
66	"4-(N,N-dimethylamino)phenylmethyl"	propyl
67	benzyl	n-butyl
68	1-pyrollylmethyl	n-butyl
69	1-pyrazolylmethyl	n-butyl
70	1-imidazolylmethyl	n-butyl
71	1-indolylmethyl	n-butyl
72	1-triazolylmethyl	n-butyl
73	1-tetrazolylmethyl	n-butyl
74	2-pyridylmethyl	n-butyl
75	3-pyridylmethyl	n-butyl

Table 3

76	4-pyridylmethyl	n-butyl
77	cyclohexylmethyl	n-butyl
78	2-naphthylmethyl	n-butyl
79	3-naphthylmethyl	n-butyl
80	2-thiophenylmethyl	n-butyl
81	4-(1-methyl)piperidinyl-	n-butyl
	methyl	
82	"(3,4-methylenedioxyphenyl)methyl"	n-butyl
	2-thienylmethyl	n-butyl
83	4-biphenylmethyl	n-butyl
84	pyrimidinylmethyl	n-butyl
85	2-benzothiazolylmethyl	n-butyl
86	2-benzothiophenylmethyl	n-butyl
87	2-thiomethylethyl	n-butyl
88	2-thiomethylmethyl	n-butyl
89	2-methylpropyl	n-butyl
90	2-methylbutyl	n-butyl
91	3-methylbutyl	n-butyl
92	cyclopropylmethyl	n-butyl
93	cyclobutylmethyl	n-butyl
94	cyclopentylmethyl	n-butyl
95	p-hydroxyphenylmethyl	n-butyl
96	p-nitrophenylmethyl	n-butyl
97	p-aminophenylmethyl	n-butyl
98	"4-(N,N-dimethylamino)phenylmethyl"	n-butyl
99	benzyl	isobutyl
100	1-pyrolylmethyl	isobutyl
101	1-pyrazolylmethyl	isobutyl
102	1-imidazolylmethyl	isobutyl
103	1-indolylmethyl	isobutyl
104	1-triazolylmethyl	isobutyl
105	1-tetrazolylmethyl	isobutyl
106	2-pyridylmethyl	isobutyl
107	3-pyridylmethyl	isobutyl
108	4-pyridylmethyl	isobutyl
109	cyclohexylmethyl	isobutyl
110	2-naphthylmethyl	isobutyl
111	3-naphthylmethyl	isobutyl
112	2-thiophenylmethyl	isobutyl
113	4-(1-methyl)piperidinyl-	isobutyl
114	methyl	
	"(3,4-methylenedioxyphenyl)methyl"	isobutyl
115	2-thienylmethyl	isobutyl
116	4-biphenylmethyl	isobutyl
117	pyrimidinylmethyl	isobutyl
118	2-benzothiazolylmethyl	isobutyl
119	2-benzothiophenylmethyl	isobutyl
120	2-thiomethylethyl	isobutyl
121	2-thiomethylmethyl	isobutyl
122	2-methylpropyl	isobutyl
123	2-methylbutyl	isobutyl
124		

Table 3

125	3-methylbutyl	isobutyl
126	cyclopropylmethyl	isobutyl
127	cyclobutylmethyl	isobutyl
128	cyclopentylmethyl	isobutyl
129	p-hydroxyphenylmethyl	isobutyl
130	p-nitrophenylmethyl	isobutyl
131	p-aminophenylmethyl	isobutyl
132	"4-(N,N-dimethylamino)phenylmethyl"	isobutyl
133	benzyl	2-butyl
134	1-pyrolylmethyl	2-butyl
135	1-pyrazolylmethyl	2-butyl
136	1-imidazolylmethyl	2-butyl
137	1-indolylmethyl	2-butyl
138	1-triazolylmethyl	2-butyl
139	1-tetrazolylmethyl	2-butyl
140	2-pyridylmethyl	2-butyl
141	3-pyridylmethyl	2-butyl
142	4-pyridylmethyl	2-butyl
143	cyclohexylmethyl	2-butyl
144	2-naphthylmethyl	2-butyl
145	3-naphthylmethyl	2-butyl
146	2-thiophenylmethyl	2-butyl
147	4-(1-methyl)piperidinyl-methyl	2-butyl
148	"(3,4-methylenedioxyphenyl)methyl"	2-butyl
149	2-thienylmethyl	2-butyl
150	4-biphenylmethyl	2-butyl
151	pyrimidinylmethyl	2-butyl
152	2-benzothiazolylmethyl	2-butyl
153	2-benzothiophenylmethyl	2-butyl
154	2-thiomethylethyl	2-butyl
155	2-thiomethylmethyl	2-butyl
156	2-methylpropyl	2-butyl
157	2-methylbutyl	2-butyl
158	3-methylbutyl	2-butyl
159	cyclopropylmethyl	2-butyl
160	cyclobutylmethyl	2-butyl
161	cyclopentylmethyl	2-butyl
162	p-hydroxyphenylmethyl	2-butyl
163	p-nitrophenylmethyl	2-butyl
164	p-aminophenylmethyl	2-butyl
165	"4-(N,N-dimethylamino)phenylmethyl"	2-butyl
166	benzyl	"3,3-dimethylallyl"
167	1-pyrolylmethyl	"3,3-dimethylallyl"
168	1-pyrazolylmethyl	"3,3-dimethylallyl"
169	1-imidazolylmethyl	"3,3-dimethylallyl"
170	1-indolylmethyl	"3,3-dimethylallyl"
171	1-triazolylmethyl	"3,3-dimethylallyl"
172	1-tetrazolylmethyl	"3,3-dimethylallyl"
173	2-pyridylmethyl	"3,3-dimethylallyl"
174	3-pyridylmethyl	"3,3-dimethylallyl"

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Table 3

175	4-pyridylmethyl	"3,3-dimethallyl"
176	cyclohexylmethyl	"3,3-dimethallyl"
177	2-naphthylmethyl	"3,3-dimethallyl"
178	3-naphthylmethyl	"3,3-dimethallyl"
179	2-thiophenylmethyl	"3,3-dimethallyl"
180	4-(1-methyl)piperidinyl-	"3,3-dimethallyl"
	methyl	
181	"(3,4-	"3,3-dimethallyl"
	methylenedioxyphenyl)methyl"	
182	2-thienylmethyl	"3,3-dimethallyl"
183	4-biphenylmethyl	"3,3-dimethallyl"
184	pyrimidinylmethyl	"3,3-dimethallyl"
185	2-benzothiazolylmethyl	"3,3-dimethallyl"
186	2-benzothiophenylmethyl	"3,3-dimethallyl"
187	2-thiomethylethyl	"3,3-dimethallyl"
188	2-thiomethylmethyl	"3,3-dimethallyl"
189	2-methylpropyl	"3,3-dimethallyl"
190	2-methylbutyl	"3,3-dimethallyl"
191	3-methylbutyl	"3,3-dimethallyl"
192	cyclopropylmethyl	"3,3-dimethallyl"
193	cyclobutylmethyl	"3,3-dimethallyl"
194	cyclopentylmethyl	"3,3-dimethallyl"
195	p-hydroxyphenylmethyl	"3,3-dimethallyl"
196	p-nitrophenylmethyl	"3,3-dimethallyl"
197	p-aminophenylmethyl	"3,3-dimethallyl"
198	"4-(N,N-	"3,3-dimethallyl"
	dimethylamino)phenylmethyl"	
199	benzyl	3-methallyl
200	1-pyrollylmethyl	3-methallyl
201	1-pyrazolylmethyl	3-methallyl
202	1-imidazolylmethyl	3-methallyl
203	1-indolylmethyl	3-methallyl
204	1-triazolylmethyl	3-methallyl
205	1-tetrazolylmethyl	3-methallyl
206	2-pyridylmethyl	3-methallyl
207	3-pyridylmethyl	3-methallyl
208	4-pyridylmethyl	3-methallyl
209	cyclohexylmethyl	3-methallyl
210	2-naphthylmethyl	3-methallyl
211	3-naphthylmethyl	3-methallyl
212	2-thiophenylmethyl	3-methallyl
213	4-(1-methyl)piperidinyl-	3-methallyl
	methyl	
214	"(3,4-	3-methallyl
	methylenedioxyphenyl)methyl"	
215	2-thienylmethyl	3-methallyl
216	4-biphenylmethyl	3-methallyl
217	pyrimidinylmethyl	3-methallyl
218	2-benzothiazolylmethyl	3-methallyl
219	2-benzothiophenylmethyl	3-methallyl
220	2-thiomethylethyl	3-methallyl
221	2-thiomethylmethyl	3-methallyl
222	2-methylpropyl	3-methallyl
223	2-methylbutyl	3-methallyl

Table 3

224	3-methylbutyl	3-methallyl
225	cyclopropylmethyl	3-methallyl
226	cyclobutylmethyl	3-methallyl
227	cyclopentylmethyl	3-methallyl
228	p-hydroxyphenylmethyl	3-methallyl
229	p-nitrophenylmethyl	3-methallyl
230	p-aminophenylmethyl	3-methallyl
231	"4-(N,N-dimethylamino)phenylmethyl"	3-methallyl
232	benzyl	2-methallyl
233	1-pyrollylmethyl	2-methallyl
234	1-pyrazolylmethyl	2-methallyl
235	1-imidazolylmethyl	2-methallyl
236	1-indolylmethyl	2-methallyl
237	1-triazolylmethyl	2-methallyl
238	1-tetrazolylmethyl	2-methallyl
239	2-pyridylmethyl	2-methallyl
240	3-pyridylmethyl	2-methallyl
241	4-pyridylmethyl	2-methallyl
242	cyclohexylmethyl	2-methallyl
243	2-naphthylmethyl	2-methallyl
244	3-naphthylmethyl	2-methallyl
245	2-thiophenylmethyl	2-methallyl
246	4-(1-methyl)piperidinyl-methyl	2-methallyl
247	"(3,4-methylenedioxyphenyl)methyl"	2-methallyl
248	2-thienylmethyl	2-methallyl
249	4-biphenylmethyl	2-methallyl
250	pyrimidinylmethyl	2-methallyl
251	2-benzothiazolylmethyl	2-methallyl
252	2-benzothiophenylmethyl	2-methallyl
253	2-thiomethylethyl	2-methallyl
254	2-thiomethylmethyl	2-methallyl
255	2-methylpropyl	2-methallyl
256	2-methylbutyl	2-methallyl
257	3-methylbutyl	2-methallyl
258	cyclopropylmethyl	2-methallyl
259	cyclobutylmethyl	2-methallyl
260	cyclopentylmethyl	2-methallyl
261	p-hydroxyphenylmethyl	2-methallyl
262	p-nitrophenylmethyl	2-methallyl
263	p-aminophenylmethyl	2-methallyl
264	"4-(N,N-dimethylamino)phenylmethyl"	2-methallyl
265	benzyl	2-propyl
266	1-pyrollylmethyl	2-propyl
267	1-pyrazolylmethyl	2-propyl
268	1-imidazolylmethyl	2-propyl
269	1-indolylmethyl	2-propyl
270	1-triazolylmethyl	2-propyl
271	1-tetrazolylmethyl	2-propyl
272	2-pyridylmethyl	2-propyl
273	3-pyridylmethyl	2-propyl

Table 3

274	4-pyridylmethyl	2-propyl
275	cyclohexylmethyl	2-propyl
276	2-naphthylmethyl	2-propyl
277	3-naphthylmethyl	2-propyl
278	2-thiophenylmethyl	2-propyl
279	4-(1-methyl)piperidinyl-	2-propyl
	methyl	
280	"(3,4-	2-propyl
	methylenedioxyphenyl)methyl"	
281	2-thienylmethyl	2-propyl
282	4-biphenylmethyl	2-propyl
283	pyrimidinylmethyl	2-propyl
284	2-benzothiazolylmethyl	2-propyl
285	2-benzothiophenylmethyl	2-propyl
286	2-thiomethylethyl	2-propyl
287	2-thiomethylmethyl	2-propyl
288	2-methylpropyl	2-propyl
289	2-methylbutyl	2-propyl
290	3-methylbutyl	2-propyl
291	cyclopropylmethyl	2-propyl
292	cyclobutylmethyl	2-propyl
293	cyclopentylmethyl	2-propyl
294	p-hydroxyphenylmethyl	2-propyl
295	p-nitrophenylmethyl	2-propyl
296	p-aminophenylmethyl	2-propyl
297	"4-(N,N-	2-propyl
	dimethylamino)phenylmethyl"	
298	benzyl	cyclopropyl
299	1-pyrollylmethyl	cyclopropyl
300	1-pyrazolylmethyl	cyclopropyl
301	1-imidazolylmethyl	cyclopropyl
302	1-indolylmethyl	cyclopropyl
303	1-triazolylmethyl	cyclopropyl
304	1-tetrazolylmethyl	cyclopropyl
305	2-pyridylmethyl	cyclopropyl
306	3-pyridylmethyl	cyclopropyl
307	4-pyridylmethyl	cyclopropyl
308	cyclohexylmethyl	cyclopropyl
309	2-naphthylmethyl	cyclopropyl
310	3-naphthylmethyl	cyclopropyl
311	2-thiophenylmethyl	cyclopropyl
312	4-(1-methyl)piperidinyl-	cyclopropyl
	methyl	
313	"(3,4-	cyclopropyl
	methylenedioxyphenyl)methyl"	
314	2-thienylmethyl	cyclopropyl
315	4-biphenylmethyl	cyclopropyl
316	pyrimidinylmethyl	cyclopropyl
317	2-benzothiazolylmethyl	cyclopropyl
318	2-benzothiophenylmethyl	cyclopropyl
319	2-thiomethylethyl	cyclopropyl
320	2-thiomethylmethyl	cyclopropyl
321	2-methylpropyl	cyclopropyl
322	2-methylbutyl	cyclopropyl

Table 3

323	3-methylbutyl	cyclopropyl
324	cyclopropylmethyl	cyclopropyl
325	cyclobutylmethyl	cyclopropyl
326	cyclopentylmethyl	cyclopropyl
327	p-hydroxyphenylmethyl	cyclopropyl
328	p-nitrophenylmethyl	cyclopropyl
329	p-aminophenylmethyl	cyclopropyl
330	"4-(N,N-dimethylamino)phenylmethyl"	cyclopropyl
331	benzyl	cyclopropylmethyl
332	1-pyrolylmethyl	cyclopropylmethyl
333	1-pyrazolylmethyl	cyclopropylmethyl
334	1-imidazolylmethyl	cyclopropylmethyl
335	1-indolylmethyl	cyclopropylmethyl
336	1-triazolylmethyl	cyclopropylmethyl
337	1-tetrazolylmethyl	cyclopropylmethyl
338	2-pyridylmethyl	cyclopropylmethyl
339	3-pyridylmethyl	cyclopropylmethyl
340	4-pyridylmethyl	cyclopropylmethyl
341	cyclohexylmethyl	cyclopropylmethyl
342	2-naphthylmethyl	cyclopropylmethyl
343	3-naphthylmethyl	cyclopropylmethyl
344	2-thiophenylmethyl	cyclopropylmethyl
345	4-(1-methyl)piperidinyl-methyl	cyclopropylmethyl
346	"(3,4-methylenedioxyphenyl)methyl"	cyclopropylmethyl
347	2-thienylmethyl	cyclopropylmethyl
348	4-biphenylmethyl	cyclopropylmethyl
349	pyrimidinylmethyl	cyclopropylmethyl
350	2-benzothiazolylmethyl	cyclopropylmethyl
351	2-benzothiophenylmethyl	cyclopropylmethyl
352	2-thiomethylethyl	cyclopropylmethyl
353	2-thiomethylmethyl	cyclopropylmethyl
354	2-methylpropyl	cyclopropylmethyl
355	2-methylbutyl	cyclopropylmethyl
356	3-methylbutyl	cyclopropylmethyl
357	cyclopropylmethyl	cyclopropylmethyl
358	cyclobutylmethyl	cyclopropylmethyl
359	cyclopentylmethyl	cyclopropylmethyl
360	p-hydroxyphenylmethyl	cyclopropylmethyl
361	p-nitrophenylmethyl	cyclopropylmethyl
362	p-aminophenylmethyl	cyclopropylmethyl
363	"4-(N,N-dimethylamino)phenylmethyl"	cyclopropylmethyl
364	benzyl	n-pentyl
365	1-pyrolylmethyl	n-pentyl
366	1-pyrazolylmethyl	n-pentyl
367	1-imidazolylmethyl	n-pentyl
368	1-indolylmethyl	n-pentyl
369	1-triazolylmethyl	n-pentyl
370	1-tetrazolylmethyl	n-pentyl
371	2-pyridylmethyl	n-pentyl
372	3-pyridylmethyl	n-pentyl

Table 3

373	4-pyridylmethyl	n-pentyl
374	cyclohexylmethyl	n-pentyl
375	2-naphthylmethyl	n-pentyl
376	3-naphthylmethyl	n-pentyl
377	2-thiophenylmethyl	n-pentyl
378	4-(1-methyl)piperidinyl-	n-pentyl
	methyl	
379	"(3,4-	n-pentyl
	methylenedioxyphenyl)methyl"	
380	2-thienylmethyl	n-pentyl
381	4-biphenylmethyl	n-pentyl
382	pyrimidinylmethyl	n-pentyl
383	2-benzothiazolylmethyl	n-pentyl
384	2-benzothiophenylmethyl	n-pentyl
385	2-thiomethylethyl	n-pentyl
386	2-thiomethylmethyl	n-pentyl
387	2-methylpropyl	n-pentyl
388	2-methylbutyl	n-pentyl
389	3-methylbutyl	n-pentyl
390	cyclopropylmethyl	n-pentyl
391	cyclobutylmethyl	n-pentyl
392	cyclopentylmethyl	n-pentyl
393	p-hydroxyphenylmethyl	n-pentyl
394	p-nitrophenylmethyl	n-pentyl
395	p-aminophenylmethyl	n-pentyl
396	"4-(N,N-	n-pentyl
	dimethylamino)phenylmethyl"	
397	benzyl	2-pentyl
398	1-pyrollylmethyl	2-pentyl
399	1-pyrazolylmethyl	2-pentyl
400	1-imidazolylmethyl	2-pentyl
401	1-indolylmethyl	2-pentyl
402	1-triazolylmethyl	2-pentyl
403	1-tetrazolylmethyl	2-pentyl
404	2-pyridylmethyl	2-pentyl
405	3-pyridylmethyl	2-pentyl
406	4-pyridylmethyl	2-pentyl
407	cyclohexylmethyl	2-pentyl
408	2-naphthylmethyl	2-pentyl
409	3-naphthylmethyl	2-pentyl
410	2-thiophenylmethyl	2-pentyl
411	4-(1-methyl)piperidinyl-	2-pentyl
	methyl	
412	"(3,4-	2-pentyl
	methylenedioxyphenyl)methyl"	
413	2-thienylmethyl	2-pentyl
414	4-biphenylmethyl	2-pentyl
415	pyrimidinylmethyl	2-pentyl
416	2-benzothiazolylmethyl	2-pentyl
417	2-benzothiophenylmethyl	2-pentyl
418	2-thiomethylethyl	2-pentyl
419	2-thiomethylmethyl	2-pentyl
420	2-methylpropyl	2-pentyl
421	2-methylbutyl	2-pentyl

Table 3

422	3-methylbutyl	2-pentyl
423	cyclopropylmethyl	2-pentyl
424	cyclobutylmethyl	2-pentyl
425	cyclopentylmethyl	2-pentyl
426	p-hydroxyphenylmethyl	2-pentyl
427	p-nitrophenylmethyl	2-pentyl
428	p-aminophenylmethyl	2-pentyl
429	"4-(N,N-dimethylamino)phenylmethyl"	2-pentyl
430	benzyl	3-pentyl
431	1-pyrollylmethyl	3-pentyl
432	1-pyrazolylmethyl	3-pentyl
433	1-imidazolylmethyl	3-pentyl
434	1-indolylmethyl	3-pentyl
435	1-triazolylmethyl	3-pentyl
436	1-tetrazolylmethyl	3-pentyl
437	2-pyridylmethyl	3-pentyl
438	3-pyridylmethyl	3-pentyl
439	4-pyridylmethyl	3-pentyl
440	cyclohexylmethyl	3-pentyl
441	2-naphthylmethyl	3-pentyl
442	3-naphthylmethyl	3-pentyl
443	2-thiophenylmethyl	3-pentyl
444	4-(1-methyl)piperidinyl-methyl	3-pentyl
445	"(3,4-methylenedioxyphenyl)methyl"	3-pentyl
446	2-thienylmethyl	3-pentyl
447	4-biphenylmethyl	3-pentyl
448	pyrimidinylmethyl	3-pentyl
449	2-benzothiazolylmethyl	3-pentyl
450	2-benzothiophenylmethyl	3-pentyl
451	2-thiomethylethyl	3-pentyl
452	2-thiomethylmethyl	3-pentyl
453	2-methylpropyl	3-pentyl
454	2-methylbutyl	3-pentyl
455	3-methylbutyl	3-pentyl
456	cyclopropylmethyl	3-pentyl
457	cyclobutylmethyl	3-pentyl
458	cyclopentylmethyl	3-pentyl
459	p-hydroxyphenylmethyl	3-pentyl
460	p-nitrophenylmethyl	3-pentyl
461	p-aminophenylmethyl	3-pentyl
462	"4-(N,N-dimethylamino)phenylmethyl"	3-pentyl
463	benzyl	3-methylbutyl
464	1-pyrollylmethyl	3-methylbutyl
465	1-pyrazolylmethyl	3-methylbutyl
466	1-imidazolylmethyl	3-methylbutyl
467	1-indolylmethyl	3-methylbutyl
468	1-triazolylmethyl	3-methylbutyl
469	1-tetrazolylmethyl	3-methylbutyl
470	2-pyridylmethyl	3-methylbutyl
471	3-pyridylmethyl	3-methylbutyl

Table 3

472	4-pyridylmethyl	3-methylbutyl
473	cyclohexylmethyl	3-methylbutyl
474	2-naphthylmethyl	3-methylbutyl
475	3-naphthylmethyl	3-methylbutyl
476	2-thiophenylmethyl	3-methylbutyl
477	4-(1-methyl)piperidinyl-	3-methylbutyl
	methyl	
478	"(3,4-	3-methylbutyl
	methylenedioxyphenyl)methyl"	
479	2-thienylmethyl	3-methylbutyl
480	4-biphenylmethyl	3-methylbutyl
481	pyrimidinylmethyl	3-methylbutyl
482	2-benzothiazolylmethyl	3-methylbutyl
483	2-benzothiophenylmethyl	3-methylbutyl
484	2-thiomethylethyl	3-methylbutyl
485	2-thiomethylmethyl	3-methylbutyl
486	2-methylpropyl	3-methylbutyl
487	2-methylbutyl	3-methylbutyl
488	3-methylbutyl	3-methylbutyl
489	cyclopropylmethyl	3-methylbutyl
490	cyclobutylmethyl	3-methylbutyl
491	cyclopentylmethyl	3-methylbutyl
492	p-hydroxyphenylmethyl	3-methylbutyl
493	p-nitrophenylmethyl	3-methylbutyl
494	p-aminophenylmethyl	3-methylbutyl
495	"4-(N,N-	3-methylbutyl
	dimethylamino)phenylmethyl"	
496	benzyl	2-methylbutyl
497	1-pyrollylmethyl	2-methylbutyl
498	1-pyrazolylmethyl	2-methylbutyl
499	1-imidazolylmethyl	2-methylbutyl
500	1-indolylmethyl	2-methylbutyl
501	1-triazolylmethyl	2-methylbutyl
502	1-tetrazolylmethyl	2-methylbutyl
503	2-pyridylmethyl	2-methylbutyl
504	3-pyridylmethyl	2-methylbutyl
505	4-pyridylmethyl	2-methylbutyl
506	cyclohexylmethyl	2-methylbutyl
507	2-naphthylmethyl	2-methylbutyl
508	3-naphthylmethyl	2-methylbutyl
509	2-thiophenylmethyl	2-methylbutyl
510	4-(1-methyl)piperidinyl-	2-methylbutyl
	methyl	
511	"(3,4-	2-methylbutyl
	methylenedioxyphenyl)methyl"	
512	2-thienylmethyl	2-methylbutyl
513	4-biphenylmethyl	2-methylbutyl
514	pyrimidinylmethyl	2-methylbutyl
515	2-benzothiazolylmethyl	2-methylbutyl
516	2-benzothiophenylmethyl	2-methylbutyl
517	2-thiomethylethyl	2-methylbutyl
518	2-thiomethylmethyl	2-methylbutyl
519	2-methylpropyl	2-methylbutyl
520	2-methylbutyl	2-methylbutyl

Table 3

521	3-methylbutyl	2-methylbutyl
522	cyclopropylmethyl	2-methylbutyl
523	cyclobutylmethyl	2-methylbutyl
524	cyclopentylmethyl	2-methylbutyl
525	p-hydroxyphenylmethyl	2-methylbutyl
526	p-nitrophenylmethyl	2-methylbutyl
527	p-aminophenylmethyl	2-methylbutyl
528	"4-(N,N-dimethylamino)phenylmethyl"	2-methylbutyl
529	benzyl	propargyl
530	1-pyrolylmethyl	propargyl
531	1-pyrazolylmethyl	propargyl
532	1-imidazolylmethyl	propargyl
533	1-indolylmethyl	propargyl
534	1-triazolylmethyl	propargyl
535	1-tetrazolylmethyl	propargyl
536	2-pyridylmethyl	propargyl
537	3-pyridylmethyl	propargyl
538	4-pyridylmethyl	propargyl
539	cyclohexylmethyl	propargyl
540	2-naphthylmethyl	propargyl
541	3-naphthylmethyl	propargyl
542	2-thiophenylmethyl	propargyl
543	4-(1-methyl)piperidinyl-methyl	propargyl
544	"(3,4-methylenedioxyphenyl)methyl"	propargyl
545	2-thienylmethyl	propargyl
546	4-biphenylmethyl	propargyl
547	pyrimidinylmethyl	propargyl
548	2-benzothiazolylmethyl	propargyl
549	2-benzothiophenylmethyl	propargyl
550	2-thiomethylethyl	propargyl
551	2-thiomethylmethyl	propargyl
552	2-methylpropyl	propargyl
553	2-methylbutyl	propargyl
554	3-methylbutyl	propargyl
555	cyclopropylmethyl	propargyl
556	cyclobutylmethyl	propargyl
557	cyclopentylmethyl	propargyl
558	p-hydroxyphenylmethyl	propargyl
559	p-nitrophenylmethyl	propargyl
560	p-aminophenylmethyl	propargyl
561	"4-(N,N-dimethylamino)phenylmethyl"	propargyl
562	benzyl	cyclobutyl
563	1-pyrolylmethyl	cyclobutyl
564	1-pyrazolylmethyl	cyclobutyl
565	1-imidazolylmethyl	cyclobutyl
566	1-indolylmethyl	cyclobutyl
567	1-triazolylmethyl	cyclobutyl
568	1-tetrazolylmethyl	cyclobutyl
569	2-pyridylmethyl	cyclobutyl
570	3-pyridylmethyl	cyclobutyl

Table 3

571	4-pyridylmethyl	cyclobutyl
572	cyclohexylmethyl	cyclobutyl
573	2-naphthylmethyl	cyclobutyl
574	3-naphthylmethyl	cyclobutyl
575	2-thiophenylmethyl	cyclobutyl
576	4-(1-methyl)piperidinyl-	cyclobutyl
	methyl	
577	"(3,4-	cyclobutyl
	methylenedioxyphenyl)methyl"	
578	2-thienylmethyl	cyclobutyl
579	4-biphenylmethyl	cyclobutyl
580	pyrimidinylmethyl	cyclobutyl
581	2-benzothiazolylmethyl	cyclobutyl
582	2-benzothiophenylmethyl	cyclobutyl
583	2-thiomethylethyl	cyclobutyl
584	2-thiomethylmethyl	cyclobutyl
585	2-methylpropyl	cyclobutyl
586	2-methylbutyl	cyclobutyl
587	3-methylbutyl	cyclobutyl
588	cyclopropylmethyl	cyclobutyl
589	cyclobutylmethyl	cyclobutyl
590	cyclopentylmethyl	cyclobutyl
591	p-hydroxyphenylmethyl	cyclobutyl
592	p-nitrophenylmethyl	cyclobutyl
593	p-aminophenylmethyl	cyclobutyl
594	"4-(N,N-	cyclobutyl
	dimethylamino)phenylmethyl"	
595	benzyl	cyclobutylmethyl
596	1-pyrollylmethyl	cyclobutylmethyl
597	1-pyrazolylmethyl	cyclobutylmethyl
598	1-imidazolylmethyl	cyclobutylmethyl
599	1-indolylmethyl	cyclobutylmethyl
600	1-triazolylmethyl	cyclobutylmethyl
601	1-tetrazolylmethyl	cyclobutylmethyl
602	2-pyridylmethyl	cyclobutylmethyl
603	3-pyridylmethyl	cyclobutylmethyl
604	4-pyridylmethyl	cyclobutylmethyl
605	cyclohexylmethyl	cyclobutylmethyl
606	2-naphthylmethyl	cyclobutylmethyl
607	3-naphthylmethyl	cyclobutylmethyl
608	2-thiophenylmethyl	cyclobutylmethyl
609	4-(1-methyl)piperidinyl-	cyclobutylmethyl
	methyl	
610	"(3,4-	cyclobutylmethyl
	methylenedioxyphenyl)methyl"	
611	2-thienylmethyl	cyclobutylmethyl
612	4-biphenylmethyl	cyclobutylmethyl
613	pyrimidinylmethyl	cyclobutylmethyl
614	2-benzothiazolylmethyl	cyclobutylmethyl
615	2-benzothiophenylmethyl	cyclobutylmethyl
616	2-thiomethylethyl	cyclobutylmethyl
617	2-thiomethylmethyl	cyclobutylmethyl
618	2-methylpropyl	cyclobutylmethyl
619	2-methylbutyl	cyclobutylmethyl

Table 3

620	3-methylbutyl	cyclobutylmethyl
621	cyclopropylmethyl	cyclobutylmethyl
622	cyclobutylmethyl	cyclobutylmethyl
623	cyclopentylmethyl	cyclobutylmethyl
624	p-hydroxyphenylmethyl	cyclobutylmethyl
625	p-nitrophenylmethyl	cyclobutylmethyl
626	p-aminophenylmethyl	cyclobutylmethyl
627	"4-(N,N-dimethylamino)phenylmethyl"	cyclobutylmethyl
628	benzyl	cyclopentyl
629	1-pyrollylmethyl	cyclopentyl
630	1-pyrazolylmethyl	cyclopentyl
631	1-imidazolylmethyl	cyclopentyl
632	1-indolylmethyl	cyclopentyl
633	1-triazolylmethyl	cyclopentyl
634	1-tetrazolylmethyl	cyclopentyl
635	2-pyridylmethyl	cyclopentyl
636	3-pyridylmethyl	cyclopentyl
637	4-pyridylmethyl	cyclopentyl
638	cyclohexylmethyl	cyclopentyl
639	2-naphthylmethyl	cyclopentyl
640	3-naphthylmethyl	cyclopentyl
641	2-thiophenylmethyl	cyclopentyl
642	4-(1-methyl)piperidinyl-methyl	cyclopentyl
643	"(3,4-methylenedioxyphenyl)methyl"	cyclopentyl
644	2-thienylmethyl	cyclopentyl
645	4-biphenylmethyl	cyclopentyl
646	pyrimidinylmethyl	cyclopentyl
647	2-benzothiazolylmethyl	cyclopentyl
648	2-benzothiophenylmethyl	cyclopentyl
649	2-thiomethylethyl	cyclopentyl
650	2-thiomethylmethyl	cyclopentyl
651	2-methylpropyl	cyclopentyl
652	2-methylbutyl	cyclopentyl
653	3-methylbutyl	cyclopentyl
654	cyclopropylmethyl	cyclopentyl
655	cyclobutylmethyl	cyclopentyl
656	cyclopentylmethyl	cyclopentyl
657	p-hydroxyphenylmethyl	cyclopentyl
658	p-nitrophenylmethyl	cyclopentyl
659	p-aminophenylmethyl	cyclopentyl
660	"4-(N,N-dimethylamino)phenylmethyl"	cyclopentyl
661	benzyl	cyclopentylmethyl
662	1-pyrollylmethyl	cyclopentylmethyl
663	1-pyrazolylmethyl	cyclopentylmethyl
664	1-imidazolylmethyl	cyclopentylmethyl
665	1-indolylmethyl	cyclopentylmethyl
666	1-triazolylmethyl	cyclopentylmethyl
667	1-tetrazolylmethyl	cyclopentylmethyl
668	2-pyridylmethyl	cyclopentylmethyl
669	3-pyridylmethyl	cyclopentylmethyl

Table 3

670	4-pyridylmethyl	cyclopentylmethyl
671	cyclohexylmethyl	cyclopentylmethyl
672	2-naphthylmethyl	cyclopentylmethyl
673	3-naphthylmethyl	cyclopentylmethyl
674	2-thiophenylmethyl	cyclopentylmethyl
675	4-(1-methyl)piperidinyl-	cyclopentylmethyl
	methyl	
676	"(3,4-	cyclopentylmethyl
	methylenedioxyphenyl)methyl"	
677	2-thienylmethyl	cyclopentylmethyl
678	4-biphenylmethyl	cyclopentylmethyl
679	pyrimidinylmethyl	cyclopentylmethyl
680	2-benzothiazolylmethyl	cyclopentylmethyl
681	2-benzothiophenylmethyl	cyclopentylmethyl
682	2-thiomethylethyl	cyclopentylmethyl
683	2-thiomethylmethyl	cyclopentylmethyl
684	2-methylpropyl	cyclopentylmethyl
685	2-methylbutyl	cyclopentylmethyl
686	3-methylbutyl	cyclopentylmethyl
687	cyclopropylmethyl	cyclopentylmethyl
688	cyclobutylmethyl	cyclopentylmethyl
689	cyclopentylmethyl	cyclopentylmethyl
690	p-hydroxyphenylmethyl	cyclopentylmethyl
691	p-nitrophenylmethyl	cyclopentylmethyl
692	p-aminophenylmethyl	cyclopentylmethyl
693	"4-(N,N-	
	dimethylamino)phenylmethyl"	
694	benzyl	1-hexyl
695	1-pyrollylmethyl	1-hexyl
696	1-pyrazolylmethyl	1-hexyl
697	1-imidazolylmethyl	1-hexyl
698	1-indolylmethyl	1-hexyl
699	1-triazolylmethyl	1-hexyl
700	1-tetrazolylmethyl	1-hexyl
701	2-pyridylmethyl	1-hexyl
702	3-pyridylmethyl	1-hexyl
703	4-pyridylmethyl	1-hexyl
704	cyclohexylmethyl	1-hexyl
705	2-naphthylmethyl	1-hexyl
706	3-naphthylmethyl	1-hexyl
707	2-thiophenylmethyl	1-hexyl
708	4-(1-methyl)piperidinyl-	1-hexyl
	methyl	
709	"(3,4-	1-hexyl
	methylenedioxyphenyl)methyl"	
710	2-thienylmethyl	1-hexyl
711	4-biphenylmethyl	1-hexyl
712	pyrimidinylmethyl	1-hexyl
713	2-benzothiazolylmethyl	1-hexyl
714	2-benzothiophenylmethyl	1-hexyl
715	2-thiomethylethyl	1-hexyl
716	2-thiomethylmethyl	1-hexyl
717	2-methylpropyl	1-hexyl
718	2-methylbutyl	1-hexyl

Table 3

719	3-methylbutyl	1-hexyl
720	cyclopropylmethyl	1-hexyl
721	cyclobutylmethyl	1-hexyl
722	cyclopentylmethyl	1-hexyl
723	p-hydroxyphenylmethyl	1-hexyl
724	p-nitrophenylmethyl	1-hexyl
725	p-aminophenylmethyl	1-hexyl
726	"4-(N,N-dimethylamino)phenylmethyl"	1-hexyl
727	benzyl	4-methylpentyl
728	1-pyrollylmethyl	4-methylpentyl
729	1-pyrazolylmethyl	4-methylpentyl
730	1-imidazolylmethyl	4-methylpentyl
731	1-indolylmethyl	4-methylpentyl
732	1-triazolylmethyl	4-methylpentyl
733	1-tetrazolylmethyl	4-methylpentyl
734	2-pyridylmethyl	4-methylpentyl
735	3-pyridylmethyl	4-methylpentyl
736	4-pyridylmethyl	4-methylpentyl
737	cyclohexylmethyl	4-methylpentyl
738	2-naphthylmethyl	4-methylpentyl
739	3-naphthylmethyl	4-methylpentyl
740	2-thiophenylmethyl	4-methylpentyl
741	4-(1-methyl)piperidinylmethyl	4-methylpentyl
742	"(3,4-methylenedioxyphenyl)methyl"	4-methylpentyl
743	2-thienylmethyl	4-methylpentyl
744	4-biphenylmethyl	4-methylpentyl
745	pyrimidinylmethyl	4-methylpentyl
746	2-benzothiazolylmethyl	4-methylpentyl
747	2-benzothiophenylmethyl	4-methylpentyl
748	2-thiomethylethyl	4-methylpentyl
749	2-thiomethylmethyl	4-methylpentyl
750	2-methylpropyl	4-methylpentyl
751	2-methylbutyl	4-methylpentyl
752	3-methylbutyl	4-methylpentyl
753	cyclopropylmethyl	4-methylpentyl
754	cyclobutylmethyl	4-methylpentyl
755	cyclopentylmethyl	4-methylpentyl
756	p-hydroxyphenylmethyl	4-methylpentyl
757	p-nitrophenylmethyl	4-methylpentyl
758	p-aminophenylmethyl	4-methylpentyl
759	"4-(N,N-dimethylamino)phenylmethyl"	4-methylpentyl
760	benzyl	3-methylpentyl
761	1-pyrollylmethyl	3-methylpentyl
762	1-pyrazolylmethyl	3-methylpentyl
763	1-imidazolylmethyl	3-methylpentyl
764	1-indolylmethyl	3-methylpentyl
765	1-triazolylmethyl	3-methylpentyl
766	1-tetrazolylmethyl	3-methylpentyl
767	2-pyridylmethyl	3-methylpentyl
768	3-pyridylmethyl	3-methylpentyl

Table 3

769	4-pyridylmethyl	3-methylpentyl
770	cyclohexylmethyl	3-methylpentyl
771	2-naphthylmethyl	3-methylpentyl
772	3-naphthylmethyl	3-methylpentyl
773	2-thiophenylmethyl	3-methylpentyl
774	4-(1-methyl)piperidinyl-	3-methylpentyl
	methyl	
775	"(3,4-	3-methylpentyl
	methylenedioxyphenyl)methyl"	
776	2-thienylmethyl	3-methylpentyl
777	4-biphenylmethyl	3-methylpentyl
778	pyrimidinylmethyl	3-methylpentyl
779	2-benzothiazolylmethyl	3-methylpentyl
780	2-benzothiophenylmethyl	3-methylpentyl
781	2-thiomethylethyl	3-methylpentyl
782	2-thiomethylmethyl	3-methylpentyl
783	2-methylpropyl	3-methylpentyl
784	2-methylbutyl	3-methylpentyl
785	3-methylbutyl	3-methylpentyl
786	cyclopropylmethyl	3-methylpentyl
787	cyclobutylmethyl	3-methylpentyl
788	cyclopentylmethyl	3-methylpentyl
789	p-hydroxyphenylmethyl	3-methylpentyl
790	p-nitrophenylmethyl	3-methylpentyl
791	p-aminophenylmethyl	3-methylpentyl
792	"4-(N,N-	3-methylpentyl
	dimethylamino)phenylmethyl"	
793	benzyl	isopropyl
794	1-pyrollylmethyl	isopropyl
795	1-pyrazolylmethyl	isopropyl
796	1-imidazolylmethyl	isopropyl
797	1-indolylmethyl	isopropyl
798	1-triazolylmethyl	isopropyl
799	1-tetrazolylmethyl	isopropyl
800	2-pyridylmethyl	isopropyl
801	3-pyridylmethyl	isopropyl
802	4-pyridylmethyl	isopropyl
803	cyclohexylmethyl	isopropyl
804	2-naphthylmethyl	isopropyl
805	3-naphthylmethyl	isopropyl
806	2-thiophenylmethyl	isopropyl
807	4-(1-methyl)piperidinyl-	isopropyl
	methyl	
808	"(3,4-	isopropyl
	methylenedioxyphenyl)methyl"	
809	2-thienylmethyl	isopropyl
810	4-biphenylmethyl	isopropyl
811	pyrimidinylmethyl	isopropyl
812	2-benzothiazolylmethyl	isopropyl
813	2-benzothiophenylmethyl	isopropyl
814	2-thiomethylethyl	isopropyl
815	2-thiomethylmethyl	isopropyl
816	2-methylpropyl	isopropyl
817	2-methylbutyl	isopropyl

Table 3

818	3-methylbutyl	isopropyl
819	cyclopropylmethyl	isopropyl
820	cyclobutylmethyl	isopropyl
821	cyclopentylmethyl	isopropyl
822	p-hydroxyphenylmethyl	isopropyl
823	p-nitrophenylmethyl	isopropyl
824	p-aminophenylmethyl	isopropyl
825	"4- (N,N-dimethylamino)phenylmethyl"	isopropyl
826	benzyl	3-methylbutyl
827	1-pyrolylmethyl	3-methylbutyl
828	1-pyrazolylmethyl	3-methylbutyl
829	1-imidazolylmethyl	3-methylbutyl
830	1-indolylmethyl	3-methylbutyl
831	1-triazolylmethyl	3-methylbutyl
832	1-tetrazolylmethyl	3-methylbutyl
833	2-pyridylmethyl	3-methylbutyl
834	3-pyridylmethyl	3-methylbutyl
835	4-pyridylmethyl	3-methylbutyl
836	cyclohexylmethyl	3-methylbutyl
837	2-naphthylmethyl	3-methylbutyl
838	3-naphthylmethyl	3-methylbutyl
839	2-thiophenylmethyl	3-methylbutyl
840	4-(1-methyl)piperidinylmethyl	3-methylbutyl
841	"(3,4-methylenedioxyphenyl)methyl"	3-methylbutyl
842	2-thienylmethyl	3-methylbutyl
843	4-biphenylmethyl	3-methylbutyl
844	pyrimidinylmethyl	3-methylbutyl
845	2-benzothiazolylmethyl	3-methylbutyl
846	2-benzothiophenylmethyl	3-methylbutyl
847	2-thiomethylethyl	3-methylbutyl
848	2-thiomethylmethyl	3-methylbutyl
849	2-methylpropyl	3-methylbutyl
850	2-methylbutyl	3-methylbutyl
851	3-methylbutyl	3-methylbutyl
852	cyclopropylmethyl	3-methylbutyl
853	cyclobutylmethyl	3-methylbutyl
854	cyclopentylmethyl	3-methylbutyl
855	p-hydroxyphenylmethyl	3-methylbutyl
856	p-nitrophenylmethyl	3-methylbutyl
857	p-aminophenylmethyl	3-methylbutyl
858	"4- (N,N-dimethylamino)phenylmethyl"	3-methylbutyl
859	benzyl	2-phenylethyl
860	1-pyrolylmethyl	2-phenylethyl
861	1-pyrazolylmethyl	2-phenylethyl
862	1-imidazolylmethyl	2-phenylethyl
863	1-indolylmethyl	2-phenylethyl
864	1-triazolylmethyl	2-phenylethyl
865	1-tetrazolylmethyl	2-phenylethyl
866	2-pyridylmethyl	2-phenylethyl
867	3-pyridylmethyl	2-phenylethyl

Table 3

868	4-pyridylmethyl	2-phenylethyl
869	cyclohexylmethyl	2-phenylethyl
870	2-naphthylmethyl	2-phenylethyl
871	3-naphthylmethyl	2-phenylethyl
872	2-thiophenylmethyl	2-phenylethyl
873	4-(1-methyl)piperidinyl-	2-phenylethyl
	methyl	
874	"(3,4-	2-phenylethyl
	methylenedioxyphenyl)methyl"	
875	2-thienylmethyl	2-phenylethyl
876	4-biphenylmethyl	2-phenylethyl
877	pyrimidinylmethyl	2-phenylethyl
878	2-benzothiazolylmethyl	2-phenylethyl
879	2-benzothiophenylmethyl	2-phenylethyl
880	2-thiomethylethyl	2-phenylethyl
881	2-thiomethylmethyl	2-phenylethyl
882	2-methylpropyl	2-phenylethyl
883	2-methylbutyl	2-phenylethyl
884	3-methylbutyl	2-phenylethyl
885	cyclopropylmethyl	2-phenylethyl
886	cyclobutylmethyl	2-phenylethyl
887	cyclopentylmethyl	2-phenylethyl
888	p-hydroxyphenylmethyl	2-phenylethyl
889	p-nitrophenylmethyl	2-phenylethyl
890	p-aminophenylmethyl	2-phenylethyl
891	"4-(N,N-	2-phenylethyl
	dimethylamino)phenylmethyl"	
892	benzyl	3-phenylpropyl
893	1-pyrollylmethyl	3-phenylpropyl
894	1-pyrazolylmethyl	3-phenylpropyl
895	1-imidazolylmethyl	3-phenylpropyl
896	1-indolylmethyl	3-phenylpropyl
897	1-triazolylmethyl	3-phenylpropyl
898	1-tetrazolylmethyl	3-phenylpropyl
899	2-pyridylmethyl	3-phenylpropyl
900	3-pyridylmethyl	3-phenylpropyl
901	4-pyridylmethyl	3-phenylpropyl
902	cyclohexylmethyl	3-phenylpropyl
903	2-naphthylmethyl	3-phenylpropyl
904	3-naphthylmethyl	3-phenylpropyl
905	2-thiophenylmethyl	3-phenylpropyl
906	4-(1-methyl)piperidinyl-	3-phenylpropyl
	methyl	
907	"(3,4-	3-phenylpropyl
	methylenedioxyphenyl)methyl"	
908	2-thienylmethyl	3-phenylpropyl
909	4-biphenylmethyl	3-phenylpropyl
910	pyrimidinylmethyl	3-phenylpropyl
911	2-benzothiazolylmethyl	3-phenylpropyl
912	2-benzothiophenylmethyl	3-phenylpropyl
913	2-thiomethylethyl	3-phenylpropyl
914	2-thiomethylmethyl	3-phenylpropyl
915	2-methylpropyl	3-phenylpropyl
916	2-methylbutyl	3-phenylpropyl

Table 3

917	3-methylbutyl	3-phenylpropyl
918	cyclopropylmethyl	3-phenylpropyl
919	cyclobutylmethyl	3-phenylpropyl
920	cyclopentylmethyl	3-phenylpropyl
921	p-hydroxyphenylmethyl	3-phenylpropyl
922	p-nitrophenylmethyl	3-phenylpropyl
923	p-aminophenylmethyl	3-phenylpropyl
924	"4-(N,N-dimethylamino)phenylmethyl"	3-phenylpropyl
925	benzyl	"2-(N,N-dimethylamino)ethyl"
926	1-pyrollylmethyl	"2-(N,N-dimethylamino)ethyl"
927	1-pyrazolylmethyl	"2-(N,N-dimethylamino)ethyl"
928	1-imidazolylmethyl	"2-(N,N-dimethylamino)ethyl"
929	1-indolylmethyl	"2-(N,N-dimethylamino)ethyl"
930	1-triazolylmethyl	"2-(N,N-dimethylamino)ethyl"
931	1-tetrazolylmethyl	"2-(N,N-dimethylamino)ethyl"
932	2-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
933	3-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
934	4-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
935	cyclohexylmethyl	"2-(N,N-dimethylamino)ethyl"
936	2-naphthylmethyl	"2-(N,N-dimethylamino)ethyl"
937	3-naphthylmethyl	"2-(N,N-dimethylamino)ethyl"
938	2-thiophenylmethyl	"2-(N,N-dimethylamino)ethyl"
939	4-(1-methyl)piperidinyl-methyl	"2-(N,N-dimethylamino)ethyl"
940	"(3,4-methylenedioxyphenyl)methyl"	"2-(N,N-dimethylamino)ethyl"
941	2-thienylmethyl	"2-(N,N-dimethylamino)ethyl"
942	4-biphenylmethyl	"2-(N,N-dimethylamino)ethyl"
943	pyrimidinylmethyl	"2-(N,N-dimethylamino)ethyl"
944	2-benzothiazolylmethyl	"2-(N,N-dimethylamino)ethyl"
945	2-benzothiophenylmethyl	"2-(N,N-dimethylamino)ethyl"
946	2-thiomethylethyl	"2-(N,N-dimethylamino)ethyl"
947	2-thiomethylmethyl	"2-(N,N-dimethylamino)ethyl"
948	2-methylpropyl	"2-(N,N-dimethylamino)ethyl"
949	2-methylbutyl	"2-(N,N-dimethylamino)ethyl"
950	3-methylbutyl	"2-(N,N-dimethylamino)ethyl"
951	cyclopropylmethyl	"2-(N,N-dimethylamino)ethyl"
952	cyclobutylmethyl	"2-(N,N-dimethylamino)ethyl"
953	cyclopentylmethyl	"2-(N,N-dimethylamino)ethyl"
954	p-hydroxyphenylmethyl	"2-(N,N-dimethylamino)ethyl"
955	p-nitrophenylmethyl	"2-(N,N-dimethylamino)ethyl"
956	p-aminophenylmethyl	"2-(N,N-dimethylamino)ethyl"
957	"4-(N,N-dimethylamino)phenylmethyl"	"2-(N,N-dimethylamino)ethyl"
958	benzyl	3-oxetanylmethyl
959	1-pyrollylmethyl	3-oxetanylmethyl
960	1-pyrazolylmethyl	3-oxetanylmethyl
961	1-imidazolylmethyl	3-oxetanylmethyl
962	1-indolylmethyl	3-oxetanylmethyl
963	1-triazolylmethyl	3-oxetanylmethyl
964	1-tetrazolylmethyl	3-oxetanylmethyl
965	2-pyridylmethyl	3-oxetanylmethyl
966	3-pyridylmethyl	3-oxetanylmethyl

Table 3

967	4-pyridylmethyl	3-oxetanylmethyl
968	cyclohexylmethyl	3-oxetanylmethyl
969	2-naphthylmethyl	3-oxetanylmethyl
970	3-naphthylmethyl	3-oxetanylmethyl
971	2-thiophenylmethyl	3-oxetanylmethyl
972	4-(1-methyl)piperidinyl- methyl	3-oxetanylmethyl
973	"(3,4- methylenedioxyphenyl)methyl"	3-oxetanylmethyl
974	2-thienylmethyl	3-oxetanylmethyl
975	4-biphenylmethyl	3-oxetanylmethyl
976	pyrimidinylmethyl	3-oxetanylmethyl
977	2-benzothiazolylmethyl	3-oxetanylmethyl
978	2-benzothiophenylmethyl	3-oxetanylmethyl
979	2-thiomethylethyl	3-oxetanylmethyl
980	2-thiomethylmethyl	3-oxetanylmethyl
981	2-methylpropyl	3-oxetanylmethyl
982	2-methylbutyl	3-oxetanylmethyl
983	3-methylbutyl	3-oxetanylmethyl
984	cyclopropylmethyl	3-oxetanylmethyl
985	cyclobutylmethyl	3-oxetanylmethyl
986	cyclopentylmethyl	3-oxetanylmethyl
987	p-hydroxyphenylmethyl	3-oxetanylmethyl
988	p-nitrophenylmethyl	3-oxetanylmethyl
989	p-aminophenylmethyl	3-oxetanylmethyl
990	"4-(N,N- dimethylamino)phenylmethyl" benzyl	3-oxetanylmethyl
991	1-pyrollylmethyl	2-tetrahydrofuranyl
992	1-pyrazolylmethyl	2-tetrahydrofuranyl
993	1-imidazolylmethyl	2-tetrahydrofuranyl
994	1-indolylmethyl	2-tetrahydrofuranyl
995	1-triazolylmethyl	2-tetrahydrofuranyl
996	1-tetrazolylmethyl	2-tetrahydrofuranyl
997	2-pyridylmethyl	2-tetrahydrofuranyl
998	3-pyridylmethyl	2-tetrahydrofuranyl
999	4-pyridylmethyl	2-tetrahydrofuranyl
1000	cyclohexylmethyl	2-tetrahydrofuranyl
1001	2-naphthylmethyl	2-tetrahydrofuranyl
1002	3-naphthylmethyl	2-tetrahydrofuranyl
1003	2-thiophenylmethyl	2-tetrahydrofuranyl
1004	4-(1-methyl)piperidinyl- methyl	2-tetrahydrofuranyl
1005	"(3,4- methylenedioxyphenyl)methyl"	2-tetrahydrofuranyl
1006	2-thienylmethyl	2-tetrahydrofuranyl
1007	4-biphenylmethyl	2-tetrahydrofuranyl
1008	pyrimidinylmethyl	2-tetrahydrofuranyl
1009	2-benzothiazolylmethyl	2-tetrahydrofuranyl
1010	2-benzothiophenylmethyl	2-tetrahydrofuranyl
1011	2-thiomethylethyl	2-tetrahydrofuranyl
1012	2-thiomethylmethyl	2-tetrahydrofuranyl
1013	2-methylpropyl	2-tetrahydrofuranyl
1014	2-methylbutyl	2-tetrahydrofuranyl
1015		

Table 3

1016	3-methylbutyl	2-tetrahydrofuranyl
1017	cyclopropylmethyl	2-tetrahydrofuranyl
1018	cyclobutylmethyl	2-tetrahydrofuranyl
1019	cyclopentylmethyl	2-tetrahydrofuranyl
1020	p-hydroxyphenylmethyl	2-tetrahydrofuranyl
1021	p-nitrophenylmethyl	2-tetrahydrofuranyl
1022	p-aminophenylmethyl	2-tetrahydrofuranyl
1023	"4-(N,N-dimethylamino)phenylmethyl"	2-tetrahydrofuranyl
1024	benzyl	2-methoxypropyl
1025	1-pyrolylmethyl	2-methoxypropyl
1026	1-pyrazolylmethyl	2-methoxypropyl
1027	1-imidazolylmethyl	2-methoxypropyl
1028	1-indolylmethyl	2-methoxypropyl
1029	1-triazolylmethyl	2-methoxypropyl
1030	1-tetrazolylmethyl	2-methoxypropyl
1031	2-pyridylmethyl	2-methoxypropyl
1032	3-pyridylmethyl	2-methoxypropyl
1033	4-pyridylmethyl	2-methoxypropyl
1034	cyclohexylmethyl	2-methoxypropyl
1035	2-naphthylmethyl	2-methoxypropyl
1036	3-naphthylmethyl	2-methoxypropyl
1037	2-thiophenylmethyl	2-methoxypropyl
1038	4-(1-methyl)piperidinyl-methyl	2-methoxypropyl
1039	"(3,4-methylenedioxyphenyl)methyl"	2-methoxypropyl
1040	2-thienylmethyl	2-methoxypropyl
1041	4-biphenylmethyl	2-methoxypropyl
1042	pyrimidinylmethyl	2-methoxypropyl
1043	2-benzothiazolylmethyl	2-methoxypropyl
1044	2-benzothienophenylmethyl	2-methoxypropyl
1045	2-thiomethylethyl	2-methoxypropyl
1046	2-thiomethylmethyl	2-methoxypropyl
1047	2-methylpropyl	2-methoxypropyl
1048	2-methylbutyl	2-methoxypropyl
1049	3-methylbutyl	2-methoxypropyl
1050	cyclopropylmethyl	2-methoxypropyl
1051	cyclobutylmethyl	2-methoxypropyl
1052	cyclopentylmethyl	2-methoxypropyl
1053	p-hydroxyphenylmethyl	2-methoxypropyl
1054	p-nitrophenylmethyl	2-methoxypropyl
1055	p-aminophenylmethyl	2-methoxypropyl
1056	"4-(N,N-dimethylamino)phenylmethyl"	2-methoxypropyl
1057	benzyl	2-ethoxyethyl
1058	1-pyrolylmethyl	2-ethoxyethyl
1059	1-pyrazolylmethyl	2-ethoxyethyl
1060	1-imidazolylmethyl	2-ethoxyethyl
1061	1-indolylmethyl	2-ethoxyethyl
1062	1-triazolylmethyl	2-ethoxyethyl
1063	1-tetrazolylmethyl	2-ethoxyethyl
1064	2-pyridylmethyl	2-ethoxyethyl
1065	3-pyridylmethyl	2-ethoxyethyl

Table 3

1066	4-pyridylmethyl	2-ethoxyethyl
1067	cyclohexylmethyl	2-ethoxyethyl
1068	2-naphthylmethyl	2-ethoxyethyl
1069	3-naphthylmethyl	2-ethoxyethyl
1070	2-thiophenylmethyl	2-ethoxyethyl
1071	4-(1-methyl)piperidinyl-	2-ethoxyethyl
	methyl	
1072	"(3,4-	2-ethoxyethyl
	methylenedioxyphenyl)methyl"	
1073	2-thienylmethyl	2-ethoxyethyl
1074	4-biphenylmethyl	2-ethoxyethyl
1075	pyrimidinylmethyl	2-ethoxyethyl
1076	2-benzothiazolylmethyl	2-ethoxyethyl
1077	2-benzothiophenylmethyl	2-ethoxyethyl
1078	2-thiomethylethyl	2-ethoxyethyl
1079	2-thiomethylmethyl	2-ethoxyethyl
1080	2-methylpropyl	2-ethoxyethyl
1081	2-methylbutyl	2-ethoxyethyl
1082	3-methylbutyl	2-ethoxyethyl
1083	cyclopropylmethyl	2-ethoxyethyl
1084	cyclobutylmethyl	2-ethoxyethyl
1085	cyclopentylmethyl	2-ethoxyethyl
1086	p-hydroxyphenylmethyl	2-ethoxyethyl
1087	p-nitrophenylmethyl	2-ethoxyethyl
1088	p-aminophenylmethyl	2-ethoxyethyl
1089	"4-(N,N-	2-ethoxyethyl
	dimethylamino)phenylmethyl"	
1090	benzyl	2-(1-pyrolyl)ethyl
1091	1-pyrollylmethyl	2-(1-pyrolyl)ethyl
1092	1-pyrazolylmethyl	2-(1-pyrolyl)ethyl
1093	1-imidazolylmethyl	2-(1-pyrolyl)ethyl
1094	1-indolylmethyl	2-(1-pyrolyl)ethyl
1095	1-triazolylmethyl	2-(1-pyrolyl)ethyl
1096	1-tetrazolylmethyl	2-(1-pyrolyl)ethyl
1097	2-pyridylmethyl	2-(1-pyrolyl)ethyl
1098	3-pyridylmethyl	2-(1-pyrolyl)ethyl
1099	4-pyridylmethyl	2-(1-pyrolyl)ethyl
1100	cyclohexylmethyl	2-(1-pyrolyl)ethyl
1101	2-naphthylmethyl	2-(1-pyrolyl)ethyl
1102	3-naphthylmethyl	2-(1-pyrolyl)ethyl
1103	2-thiophenylmethyl	2-(1-pyrolyl)ethyl
1104	4-(1-methyl)piperidinyl-	2-(1-pyrolyl)ethyl
	methyl	
1105	"(3,4-	2-(1-pyrolyl)ethyl
	methylenedioxyphenyl)methyl"	
1106	2-thienylmethyl	2-(1-pyrolyl)ethyl
1107	4-biphenylmethyl	2-(1-pyrolyl)ethyl
1108	pyrimidinylmethyl	2-(1-pyrolyl)ethyl
1109	2-benzothiazolylmethyl	2-(1-pyrolyl)ethyl
1110	2-benzothiophenylmethyl	2-(1-pyrolyl)ethyl
1111	2-thiomethylethyl	2-(1-pyrolyl)ethyl
1112	2-thiomethylmethyl	2-(1-pyrolyl)ethyl
1113	2-methylpropyl	2-(1-pyrolyl)ethyl
1114	2-methylbutyl	2-(1-pyrolyl)ethyl

Table 3

1115	3-methylbutyl	2-(1-pyrolyl)ethyl
1116	cyclopropylmethyl	2-(1-pyrolyl)ethyl
1117	cyclobutylmethyl	2-(1-pyrolyl)ethyl
1118	cyclopentylmethyl	2-(1-pyrolyl)ethyl
1119	p-hydroxyphenylmethyl	2-(1-pyrolyl)ethyl
1120	p-nitrophenylmethyl	2-(1-pyrolyl)ethyl
1121	p-aminophenylmethyl	2-(1-pyrolyl)ethyl
1122	"4-(N,N-dimethylamino)phenylmethyl"	2-(1-pyrolyl)ethyl
1123	benzyl	2-(1-imidazolyl)ethyl
1124	1-pyrolylmethyl	2-(1-imidazolyl)ethyl
1125	1-pyrazolylmethyl	2-(1-imidazolyl)ethyl
1126	1-imidazolylmethyl	2-(1-imidazolyl)ethyl
1127	1-indolylmethyl	2-(1-imidazolyl)ethyl
1128	1-triazolylmethyl	2-(1-imidazolyl)ethyl
1129	1-tetrazolylmethyl	2-(1-imidazolyl)ethyl
1130	2-pyridylmethyl	2-(1-imidazolyl)ethyl
1131	3-pyridylmethyl	2-(1-imidazolyl)ethyl
1132	4-pyridylmethyl	2-(1-imidazolyl)ethyl
1133	cyclohexylmethyl	2-(1-imidazolyl)ethyl
1134	2-naphthylmethyl	2-(1-imidazolyl)ethyl
1135	3-naphthylmethyl	2-(1-imidazolyl)ethyl
1136	2-thiophenylmethyl	2-(1-imidazolyl)ethyl
1137	4-(1-methyl)piperidinylmethyl	2-(1-imidazolyl)ethyl
1138	"(3,4-methylenedioxyphenyl)methyl"	2-(1-imidazolyl)ethyl
1139	2-thienylmethyl	2-(1-imidazolyl)ethyl
1140	4-biphenylmethyl	2-(1-imidazolyl)ethyl
1141	pyrimidinylmethyl	2-(1-imidazolyl)ethyl
1142	2-benzothiazolylmethyl	2-(1-imidazolyl)ethyl
1143	2-benzothiophenylmethyl	2-(1-imidazolyl)ethyl
1144	2-thiomethylethyl	2-(1-imidazolyl)ethyl
1145	2-thiomethylmethyl	2-(1-imidazolyl)ethyl
1146	2-methylpropyl	2-(1-imidazolyl)ethyl
1147	2-methylbutyl	2-(1-imidazolyl)ethyl
1148	3-methylbutyl	2-(1-imidazolyl)ethyl
1149	cyclopropylmethyl	2-(1-imidazolyl)ethyl
1150	cyclobutylmethyl	2-(1-imidazolyl)ethyl
1151	cyclopentylmethyl	2-(1-imidazolyl)ethyl
1152	p-hydroxyphenylmethyl	2-(1-imidazolyl)ethyl
1153	p-nitrophenylmethyl	2-(1-imidazolyl)ethyl
1154	p-aminophenylmethyl	2-(1-imidazolyl)ethyl
1155	"4-(N,N-dimethylamino)phenylmethyl"	2-(1-imidazolyl)ethyl
1156	benzyl	2-pyridylmethyl
1157	1-pyrolylmethyl	2-pyridylmethyl
1158	1-pyrazolylmethyl	2-pyridylmethyl
1159	1-imidazolylmethyl	2-pyridylmethyl
1160	1-indolylmethyl	2-pyridylmethyl
1161	1-triazolylmethyl	2-pyridylmethyl
1162	1-tetrazolylmethyl	2-pyridylmethyl
1163	2-pyridylmethyl	2-pyridylmethyl
1164	3-pyridylmethyl	2-pyridylmethyl

Table 3

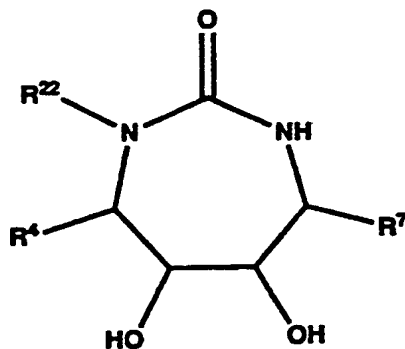
1165	4-pyridylmethyl	2-pyridylmethyl
1166	cyclohexylmethyl	2-pyridylmethyl
1167	2-naphthylmethyl	2-pyridylmethyl
1168	3-naphthylmethyl	2-pyridylmethyl
1169	2-thiophenylmethyl	2-pyridylmethyl
1170	4-(1-methyl)piperidinyl-	2-pyridylmethyl
	methyl	
1171	"(3,4-	2-pyridylmethyl
	methylenedioxyphenyl)methyl"	
1172	2-thienylmethyl	2-pyridylmethyl
1173	4-biphenylmethyl	2-pyridylmethyl
1174	pyrimidinylmethyl	2-pyridylmethyl
1175	2-benzothiazolylmethyl	2-pyridylmethyl
1176	2-benzothiophenylmethyl	2-pyridylmethyl
1177	2-thiomethylethyl	2-pyridylmethyl
1178	2-thiomethylmethyl	2-pyridylmethyl
1179	2-methylpropyl	2-pyridylmethyl
1180	2-methylbutyl	2-pyridylmethyl
1181	3-methylbutyl	2-pyridylmethyl
1182	cyclopropylmethyl	2-pyridylmethyl
1183	cyclobutylmethyl	2-pyridylmethyl
1184	cyclopentylmethyl	2-pyridylmethyl
1185	p-hydroxyphenylmethyl	2-pyridylmethyl
1186	p-nitrophenylmethyl	2-pyridylmethyl
1187	p-aminophenylmethyl	2-pyridylmethyl
1188	"4-(N,N-	2-pyridylmethyl
	dimethylamino)phenylmethyl"	
1189	benzyl	2-thiomethylethyl
1190	1-pyrolylmethyl	2-thiomethylethyl
1191	1-pyrazolylmethyl	2-thiomethylethyl
1192	1-imidazolylmethyl	2-thiomethylethyl
1193	1-indolylmethyl	2-thiomethylethyl
1194	1-triazolylmethyl	2-thiomethylethyl
1195	1-tetrazolylmethyl	2-thiomethylethyl
1196	2-pyridylmethyl	2-thiomethylethyl
1197	3-pyridylmethyl	2-thiomethylethyl
1198	4-pyridylmethyl	2-thiomethylethyl
1199	cyclohexylmethyl	2-thiomethylethyl
1200	2-naphthylmethyl	2-thiomethylethyl
1201	3-naphthylmethyl	2-thiomethylethyl
1202	2-thiophenylmethyl	2-thiomethylethyl
1203	4-(1-methyl)piperidinyl-	2-thiomethylethyl
	methyl	
1204	"(3,4-	2-thiomethylethyl
	methylenedioxyphenyl)methyl"	
1205	2-thienylmethyl	2-thiomethylethyl
1206	4-biphenylmethyl	2-thiomethylethyl
1207	pyrimidinylmethyl	2-thiomethylethyl
1208	2-benzothiazolylmethyl	2-thiomethylethyl
1209	2-benzothiophenylmethyl	2-thiomethylethyl
1210	2-thiomethylethyl	2-thiomethylethyl
1211	2-thiomethylmethyl	2-thiomethylethyl
1212	2-methylpropyl	2-thiomethylethyl
1213	2-methylbutyl	2-thiomethylethyl

Table 3

1214	3-methylbutyl	2-thiomethylethyl
1215	cyclopropylmethyl	2-thiomethylethyl
1216	cyclobutylmethyl	2-thiomethylethyl
1217	cyclopentylmethyl	2-thiomethylethyl
1218	p-hydroxyphenylmethyl	2-thiomethylethyl
1219	p-nitrophenylmethyl	2-thiomethylethyl
1220	p-aminophenylmethyl	2-thiomethylethyl
1221	"4-(N,N-dimethylamino)phenylmethyl"	2-thiomethylethyl

Table 4

TABLE 4



$$R^4 = R^7; R^{22}$$

Example $R^4 = R^7$
Number

 R^{22}

1222	benzyl	allyl
1223	1-pyrrolylmethyl	allyl
1224	1-pyrazolylmethyl	allyl
1225	1-imidazolylmethyl	allyl
1226	1-indolylmethyl	allyl
1227	1-triazolylmethyl	allyl
1228	1-tetrazolylmethyl	allyl
1229	2-pyridylmethyl	allyl
1230	3-pyridylmethyl	allyl
1231	4-pyridylmethyl	allyl
1232	cyclohexylmethyl	allyl
1233	2-naphthylmethyl	allyl
1234	3-naphthylmethyl	allyl
1235	2-thiophenylmethyl	allyl
1236	4-(1-methyl)piperidinyl- methyl	allyl
1237	"(3,4- methylenedioxyphenyl)methyl"	allyl
1238	2-thienylmethyl	allyl
1239	4-biphenylmethyl	allyl
1240	pyrimidinylmethyl	allyl
1241	2-benzothiazolylmethyl	allyl
1242	2-benzothiophenylmethyl	allyl
1243	2-thiomethylethyl	allyl
1244	2-thiomethylmethyl	allyl
1245	2-methylpropyl	allyl
1246	2-methylbutyl	allyl
1247	3-methylbutyl	allyl
1248	cyclopropylmethyl	allyl
1249	cyclobutylmethyl	allyl

Table 4

1250	cyclopentylmethyl	allyl
1251	p-hydroxyphenylmethyl	allyl
1252	p-nitrophenylmethyl	allyl
1253	p-aminophenylmethyl	allyl
1254	"4-(N,N-dimethylamino)phenylmethyl"	allyl
1255	benzyl	propyl
1256	1-pyrollylmethyl	propyl
1257	1-pyrazolylmethyl	propyl
1258	1-imidazolylmethyl	propyl
1259	1-indolylmethyl	propyl
1260	1-triazolylmethyl	propyl
1261	1-tetrazolylmethyl	propyl
1262	2-pyridylmethyl	propyl
1263	3-pyridylmethyl	propyl
1264	4-pyridylmethyl	propyl
1265	cyclohexylmethyl	propyl
1266	2-naphthylmethyl	propyl
1267	3-naphthylmethyl	propyl
1268	2-thiophenylmethyl	propyl
1269	4-(1-methyl)piperidinyl-methyl	propyl
1270	"(3,4-methylenedioxyphenyl)methyl"	propyl
1271	2-thienylmethyl	propyl
1272	4-biphenylmethyl	propyl
1273	pyrimidinylmethyl	propyl
1274	2-benzothiazolylmethyl	propyl
1275	2-benzothiophenylmethyl	propyl
1276	2-thiomethylethyl	propyl
1277	2-thiomethylmethyl	propyl
1278	2-methylpropyl	propyl
1279	2-methylbutyl	propyl
1280	3-methylbutyl	propyl
1281	cyclopropylmethyl	propyl
1282	cyclobutylmethyl	propyl
1283	cyclopentylmethyl	propyl
1284	p-hydroxyphenylmethyl	propyl
1285	p-nitrophenylmethyl	propyl
1286	p-aminophenylmethyl	propyl
1287	"4-(N,N-dimethylamino)phenylmethyl"	propyl
1288	benzyl	n-butyl
1289	1-pyrollylmethyl	n-butyl
1290	1-pyrazolylmethyl	n-butyl
1291	1-imidazolylmethyl	n-butyl
1292	1-indolylmethyl	n-butyl
1293	1-triazolylmethyl	n-butyl
1294	1-tetrazolylmethyl	n-butyl
1295	2-pyridylmethyl	n-butyl
1296	3-pyridylmethyl	n-butyl
1297	4-pyridylmethyl	n-butyl
1298	cyclohexylmethyl	n-butyl
1299	2-naphthylmethyl	n-butyl
1300	3-naphthylmethyl	n-butyl
1301	2-thiophenylmethyl	n-butyl

Table 4

1302	4-(1-methyl)piperidinyl-methyl	n-butyl
1303	"(3,4-methylenedioxyphenyl)methyl"	n-butyl
1304	2-thienylmethyl	n-butyl
1305	4-biphenylmethyl	n-butyl
1306	pyrimidinylmethyl	n-butyl
1307	2-benzothiazolylmethyl	n-butyl
1308	2-benzothiophenylmethyl	n-butyl
1309	2-thiomethylethyl	n-butyl
1310	2-thiomethylmethyl	n-butyl
1311	2-methylpropyl	n-butyl
1312	2-methylbutyl	n-butyl
1313	3-methylbutyl	n-butyl
1314	cyclopropylmethyl	n-butyl
1315	cyclobutylmethyl	n-butyl
1316	cyclopentylmethyl	n-butyl
1317	p-hydroxyphenylmethyl	n-butyl
1318	p-nitrophenylmethyl	n-butyl
1319	p-aminophenylmethyl	n-butyl
1320	"4-(N,N-dimethylamino)phenylmethyl"	isobutyl
1321	benzyl	isobutyl
1322	1-pyrollylmethyl	isobutyl
1323	1-pyrazolylmethyl	isobutyl
1324	1-imidazolylmethyl	isobutyl
1325	1-indolylmethyl	isobutyl
1326	1-triazolylmethyl	isobutyl
1327	1-tetrazolylmethyl	isobutyl
1328	2-pyridylmethyl	isobutyl
1329	3-pyridylmethyl	isobutyl
1330	4-pyridylmethyl	isobutyl
1331	cyclohexylmethyl	isobutyl
1332	2-naphthylmethyl	isobutyl
1333	3-naphthylmethyl	isobutyl
1334	2-thiophenylmethyl	isobutyl
1335	4-(1-methyl)piperidinyl-methyl	isobutyl
1336	"(3,4-methylenedioxyphenyl)methyl"	isobutyl
1337	2-thienylmethyl	isobutyl
1338	4-biphenylmethyl	isobutyl
1339	pyrimidinylmethyl	isobutyl
1340	2-benzothiazolylmethyl	isobutyl
1341	2-benzothiophenylmethyl	isobutyl
1342	2-thiomethylethyl	isobutyl
1343	2-thiomethylmethyl	isobutyl
1344	2-methylpropyl	isobutyl
1345	2-methylbutyl	isobutyl
1346	3-methylbutyl	isobutyl
1347	cyclopropylmethyl	isobutyl
1348	cyclobutylmethyl	isobutyl
1349	cyclopentylmethyl	isobutyl
1350	p-hydroxyphenylmethyl	isobutyl
1351	p-nitrophenylmethyl	isobutyl
1352	p-aminophenylmethyl	isobutyl

Table 4

1353	"4-(N,N-dimethylamino)phenylmethyl"	isobutyl
1354	benzyl	2-butyl
1355	1-pyrolylmethyl	2-butyl
1356	1-pyrazolylmethyl	2-butyl
1357	1-imidazolylmethyl	2-butyl
1358	1-indolylmethyl	2-butyl
1359	1-triazolylmethyl	2-butyl
1360	1-tetrazolylmethyl	2-butyl
1361	2-pyridylmethyl	2-butyl
1362	3-pyridylmethyl	2-butyl
1363	4-pyridylmethyl	2-butyl
1364	cyclohexylmethyl	2-butyl
1365	2-naphthylmethyl	2-butyl
1366	3-naphthylmethyl	2-butyl
1367	2-thiophenylmethyl	2-butyl
1368	4-(1-methyl)piperidinyl-methyl	2-butyl
1369	"(3,4-methylenedioxyphenyl)methyl"	2-butyl
1370	2-thienylmethyl	2-butyl
1371	4-biphenylmethyl	2-butyl
1372	pyrimidinylmethyl	2-butyl
1373	2-benzothiazolylmethyl	2-butyl
1374	2-benzothiophenylmethyl	2-butyl
1375	2-thiomethylethyl	2-butyl
1376	2-thiomethylmethyl	2-butyl
1377	2-methylpropyl	2-butyl
1378	2-methylbutyl	2-butyl
1379	3-methylbutyl	2-butyl
1380	cyclopropylmethyl	2-butyl
1381	cyclobutylmethyl	2-butyl
1382	cyclopentylmethyl	2-butyl
1383	p-hydroxyphenylmethyl	2-butyl
1384	p-nitrophenylmethyl	2-butyl
1385	p-aminophenylmethyl	2-butyl
1386	"4-(N,N-dimethylamino)phenylmethyl"	
1387	benzyl	"3,3-dimethallyl"
1388	1-pyrolylmethyl	"3,3-dimethallyl"
1389	1-pyrazolylmethyl	"3,3-dimethallyl"
1390	1-imidazolylmethyl	"3,3-dimethallyl"
1391	1-indolylmethyl	"3,3-dimethallyl"
1392	1-triazolylmethyl	"3,3-dimethallyl"
1393	1-tetrazolylmethyl	"3,3-dimethallyl"
1394	2-pyridylmethyl	"3,3-dimethallyl"
1395	3-pyridylmethyl	"3,3-dimethallyl"
1396	4-pyridylmethyl	"3,3-dimethallyl"
1397	cyclohexylmethyl	"3,3-dimethallyl"
1398	2-naphthylmethyl	"3,3-dimethallyl"
1399	3-naphthylmethyl	"3,3-dimethallyl"
1400	2-thiophenylmethyl	"3,3-dimethallyl"
1401	4-(1-methyl)piperidinyl-methyl	"3,3-dimethallyl"
1402	"(3,4-methylenedioxyphenyl)methyl"	"3,3-dimethallyl"
1403	2-thienylmethyl	"3,3-dimethallyl"

Table 4

1404	4-biphenylmethyl	"3,3-dimethallyl"
1405	pyrimidinylmethyl	"3,3-dimethallyl"
1406	2-benzothiazolylmethyl	"3,3-dimethallyl"
1407	2-benzothiophenylmethyl	"3,3-dimethallyl"
1408	2-thiomethylethyl	"3,3-dimethallyl"
1409	2-thiomethylmethyl	"3,3-dimethallyl"
1410	2-methylpropyl	"3,3-dimethallyl"
1411	2-methylbutyl	"3,3-dimethallyl"
1412	3-methylbutyl	"3,3-dimethallyl"
1413	cyclopropylmethyl	"3,3-dimethallyl"
1414	cyclobutylmethyl	"3,3-dimethallyl"
1415	cyclopentylmethyl	"3,3-dimethallyl"
1416	p-hydroxyphenylmethyl	"3,3-dimethallyl"
1417	p-nitrophenylmethyl	"3,3-dimethallyl"
1418	p-aminophenylmethyl	"3,3-dimethallyl"
1419	"4-(N,N-dimethylamino)phenylmethyl"	"3,3-dimethallyl"
1420	benzyl	3-methallyl
1421	1-pyrollylmethyl	3-methallyl
1422	1-pyrazolylmethyl	3-methallyl
1423	1-imidazolylmethyl	3-methallyl
1424	1-indolylmethyl	3-methallyl
1425	1-triazolylmethyl	3-methallyl
1426	1-tetrazolylmethyl	3-methallyl
1427	2-pyridylmethyl	3-methallyl
1428	3-pyridylmethyl	3-methallyl
1429	4-pyridylmethyl	3-methallyl
1430	cyclohexylmethyl	3-methallyl
1431	2-naphthylmethyl	3-methallyl
1432	3-naphthylmethyl	3-methallyl
1433	2-thiophenylmethyl	3-methallyl
1434	4-(1-methyl)piperidinyl-methyl	3-methallyl
1435	"(3,4-methylenedioxyphenyl)methyl"	3-methallyl
1436	2-thienylmethyl	3-methallyl
1437	4-biphenylmethyl	3-methallyl
1438	pyrimidinylmethyl	3-methallyl
1439	2-benzothiazolylmethyl	3-methallyl
1440	2-benzothiophenylmethyl	3-methallyl
1441	2-thiomethylethyl	3-methallyl
1442	2-thiomethylmethyl	3-methallyl
1443	2-methylpropyl	3-methallyl
1444	2-methylbutyl	3-methallyl
1445	3-methylbutyl	3-methallyl
1446	cyclopropylmethyl	3-methallyl
1447	cyclobutylmethyl	3-methallyl
1448	cyclopentylmethyl	3-methallyl
1449	p-hydroxyphenylmethyl	3-methallyl
1450	p-nitrophenylmethyl	3-methallyl
1451	p-aminophenylmethyl	3-methallyl
1452	"4-(N,N-dimethylamino)phenylmethyl"	3-methallyl
1453	benzyl	2-methallyl
1454	1-pyrollylmethyl	2-methallyl
1455	1-pyrazolylmethyl	2-methallyl
1456	1-imidazolylmethyl	2-methallyl

Table 4

1457	1-indolylmethyl	2-methallyl
1458	1-triazolylmethyl	2-methallyl
1459	1-tetrazolylmethyl	2-methallyl
1460	2-pyridylmethyl	2-methallyl
1461	3-pyridylmethyl	2-methallyl
1462	4-pyridylmethyl	2-methallyl
1463	cyclohexylmethyl	2-methallyl
1464	2-naphthylmethyl	2-methallyl
1465	3-naphthylmethyl	2-methallyl
1466	2-thiophenylmethyl	2-methallyl
1467	4-(1-methyl)piperidinyl-methyl	2-methallyl
1468	"(3,4-methylenedioxyphenyl)methyl"	2-methallyl
1469	2-thienylmethyl	2-methallyl
1470	4-biphenylmethyl	2-methallyl
1471	pyrimidinylmethyl	2-methallyl
1472	2-benzothiazolylmethyl	2-methallyl
1473	2-benzothiophenylmethyl	2-methallyl
1474	2-thiomethylethyl	2-methallyl
1475	2-thiomethylmethyl	2-methallyl
1476	2-methylpropyl	2-methallyl
1477	2-methylbutyl	2-methallyl
1478	3-methylbutyl	2-methallyl
1479	cyclopropylmethyl	2-methallyl
1480	cyclobutylmethyl	2-methallyl
1481	cyclopentylmethyl	2-methallyl
1482	p-hydroxyphenylmethyl	2-methallyl
1483	p-nitrophenylmethyl	2-methallyl
1484	p-aminophenylmethyl	2-methallyl
1485	"4-(N,N-dimethylamino)phenylmethyl"	2-methallyl
1486	benzyl	2-propyl
1487	1-pyrollylmethyl	2-propyl
1488	1-pyrazolylmethyl	2-propyl
1489	1-imidazolylmethyl	2-propyl
1490	1-indolylmethyl	2-propyl
1491	1-triazolylmethyl	2-propyl
1492	1-tetrazolylmethyl	2-propyl
1493	2-pyridylmethyl	2-propyl
1494	3-pyridylmethyl	2-propyl
1495	4-pyridylmethyl	2-propyl
1496	cyclohexylmethyl	2-propyl
1497	2-naphthylmethyl	2-propyl
1498	3-naphthylmethyl	2-propyl
1499	2-thiophenylmethyl	2-propyl
1500	4-(1-methyl)piperidinyl-methyl	2-propyl
1501	"(3,4-methylenedioxyphenyl)methyl"	2-propyl
1502	2-thienylmethyl	2-propyl
1503	4-biphenylmethyl	2-propyl
1504	pyrimidinylmethyl	2-propyl
1505	2-benzothiazolylmethyl	2-propyl
1506	2-benzothiophenylmethyl	2-propyl
1507	2-thiomethylethyl	2-propyl
1508	2-thiomethylmethyl	2-propyl

Table 4

1509	2-methylpropyl	2-propyl
1510	2-methylbutyl	2-propyl
1511	3-methylbutyl	2-propyl
1512	cyclopropylmethyl	2-propyl
1513	cyclobutylmethyl	2-propyl
1514	cyclopentylmethyl	2-propyl
1515	p-hydroxyphenylmethyl	2-propyl
1516	p-nitrophenylmethyl	2-propyl
1517	p-aminophenylmethyl	2-propyl
1518	"4- (N,N-dimethylamino) phenylmethyl"	2-propyl
1519	benzyl	cyclopropyl
1520	1-pyrollylmethyl	cyclopropyl
1521	1-pyrazolylmethyl	cyclopropyl
1522	1-imidazolylmethyl	cyclopropyl
1523	1-indolylmethyl	cyclopropyl
1524	1-triazolylmethyl	cyclopropyl
1525	1-tetrazolylmethyl	cyclopropyl
1526	2-pyridylmethyl	cyclopropyl
1527	3-pyridylmethyl	cyclopropyl
1528	4-pyridylmethyl	cyclopropyl
1529	cyclohexylmethyl	cyclopropyl
1530	2-naphthylmethyl	cyclopropyl
1531	3-naphthylmethyl	cyclopropyl
1532	2-thiophenylmethyl	cyclopropyl
1533	4- (1-methyl) piperidinyl-methyl	cyclopropyl
1534	" (3,4-methylenedioxyphenyl) methyl"	cyclopropyl
1535	2-thienylmethyl	cyclopropyl
1536	4-biphenylmethyl	cyclopropyl
1537	pyrimidinylmethyl	cyclopropyl
1538	2-benzothiazolylmethyl	cyclopropyl
1539	2-benzothiophenylmethyl	cyclopropyl
1540	2-thiomethylethyl	cyclopropyl
1541	2-thiomethylmethyl	cyclopropyl
1542	2-methylpropyl	cyclopropyl
1543	2-methylbutyl	cyclopropyl
1544	3-methylbutyl	cyclopropyl
1545	cyclopropylmethyl	cyclopropyl
1546	cyclobutylmethyl	cyclopropyl
1547	cyclopentylmethyl	cyclopropyl
1548	p-hydroxyphenylmethyl	cyclopropyl
1549	p-nitrophenylmethyl	cyclopropyl
1550	p-aminophenylmethyl	cyclopropyl
1551	"4- (N,N-dimethylamino) phenylmethyl"	cyclopropyl
1552	benzyl	cyclopropylmethyl
1553	1-pyrollylmethyl	cyclopropylmethyl
1554	1-pyrazolylmethyl	cyclopropylmethyl
1555	1-imidazolylmethyl	cyclopropylmethyl
1556	1-indolylmethyl	cyclopropylmethyl
1557	1-triazolylmethyl	cyclopropylmethyl
1558	1-tetrazolylmethyl	cyclopropylmethyl
1559	2-pyridylmethyl	cyclopropylmethyl
1560	3-pyridylmethyl	cyclopropylmethyl
1561	4-pyridylmethyl	cyclopropylmethyl

Table 4

1562	cyclohexylmethyl	cyclopropylmethyl
1563	2-naphthylmethyl	cyclopropylmethyl
1564	3-naphthylmethyl	cyclopropylmethyl
1565	2-thiophenylmethyl	cyclopropylmethyl
1566	4-(1-methyl)piperidinyl-	cyclopropylmethyl
	methyl	
1567	"(3,4-	cyclopropylmethyl
	methylenedioxyphenyl)methyl"	
1568	2-thienylmethyl	cyclopropylmethyl
1569	4-biphenylmethyl	cyclopropylmethyl
1570	pyrimidinylmethyl	cyclopropylmethyl
1571	2-benzothiazolylmethyl	cyclopropylmethyl
1572	2-benzothiophenylmethyl	cyclopropylmethyl
1573	2-thiomethylethyl	cyclopropylmethyl
1574	2-thiomethylmethyl	cyclopropylmethyl
1575	2-methylpropyl	cyclopropylmethyl
1576	2-methylbutyl	cyclopropylmethyl
1577	3-methylbutyl	cyclopropylmethyl
1578	cyclopropylmethyl	cyclopropylmethyl
1579	cyclobutylmethyl	cyclopropylmethyl
1580	cyclopentylmethyl	cyclopropylmethyl
1581	p-hydroxyphenylmethyl	cyclopropylmethyl
1582	p-nitrophenylmethyl	cyclopropylmethyl
1583	p-aminophenylmethyl	cyclopropylmethyl
1584	"4-(N,N-	cyclopropylmethyl
	dimethylamino)phenylmethyl"	
1585	benzyl	n-pentyl
1586	1-pyrollylmethyl	n-pentyl
1587	1-pyrazolylmethyl	n-pentyl
1588	1-imidazolylmethyl	n-pentyl
1589	1-indolylmethyl	n-pentyl
1590	1-triazolylmethyl	n-pentyl
1591	1-tetrazolylmethyl	n-pentyl
1592	2-pyridylmethyl	n-pentyl
1593	3-pyridylmethyl	n-pentyl
1594	4-pyridylmethyl	n-pentyl
1595	cyclohexylmethyl	n-pentyl
1596	2-naphthylmethyl	n-pentyl
1597	3-naphthylmethyl	n-pentyl
1598	2-thiophenylmethyl	n-pentyl
1599	4-(1-methyl)piperidinyl-	n-pentyl
	methyl	
1600	"(3,4-	n-pentyl
	methylenedioxyphenyl)methyl"	
1601	2-thienylmethyl	n-pentyl
1602	4-biphenylmethyl	n-pentyl
1603	pyrimidinylmethyl	n-pentyl
1604	2-benzothiazolylmethyl	n-pentyl
1605	2-benzothiophenylmethyl	n-pentyl
1606	2-thiomethylethyl	n-pentyl
1607	2-thiomethylmethyl	n-pentyl
1608	2-methylpropyl	n-pentyl
1609	2-methylbutyl	n-pentyl
1610	3-methylbutyl	n-pentyl
1611	cyclopropylmethyl	n-pentyl
1612	cyclobutylmethyl	n-pentyl
1613	cyclopentylmethyl	n-pentyl

Table 4

1614	p-hydroxyphenylmethyl	n-pentyl
1615	p-nitrophenylmethyl	n-pentyl
1616	p-aminophenylmethyl	n-pentyl
1617	"4-(N,N-dimethylamino) phenylmethyl"	n-pentyl
1618	benzyl	2-pentyl
1619	1-pyrollylmethyl	2-pentyl
1620	1-pyrazolylmethyl	2-pentyl
1621	1-imidazolylmethyl	2-pentyl
1622	1-indolylmethyl	2-pentyl
1623	1-triazolylmethyl	2-pentyl
1624	1-tetrazolylmethyl	2-pentyl
1625	2-pyridylmethyl	2-pentyl
1626	3-pyridylmethyl	2-pentyl
1627	4-pyridylmethyl	2-pentyl
1628	cyclohexylmethyl	2-pentyl
1629	2-naphthylmethyl	2-pentyl
1630	3-naphthylmethyl	2-pentyl
1631	2-thiophenylmethyl	2-pentyl
1632	4-(1-methyl)piperidinyl-methyl	2-pentyl
1633	"(3,4-methylenedioxyphenyl)methyl"	2-pentyl
1634	2-thienylmethyl	2-pentyl
1635	4-biphenylmethyl	2-pentyl
1636	pyrimidinylmethyl	2-pentyl
1637	2-benzothiazolylmethyl	2-pentyl
1638	2-benzothiophenylmethyl	2-pentyl
1639	2-thiomethylethyl	2-pentyl
1640	2-thiomethylmethyl	2-pentyl
1641	2-methylpropyl	2-pentyl
1642	2-methylbutyl	2-pentyl
1643	3-methylbutyl	2-pentyl
1644	cyclopropylmethyl	2-pentyl
1645	cyclobutylmethyl	2-pentyl
1646	cyclopentylmethyl	2-pentyl
1647	p-hydroxyphenylmethyl	2-pentyl
1648	p-nitrophenylmethyl	2-pentyl
1649	p-aminophenylmethyl	2-pentyl
1650	"4-(N,N-dimethylamino) phenylmethyl"	3-pentyl
1651	benzyl	3-pentyl
1652	1-pyrollylmethyl	3-pentyl
1653	1-pyrazolylmethyl	3-pentyl
1654	1-imidazolylmethyl	3-pentyl
1655	1-indolylmethyl	3-pentyl
1656	1-triazolylmethyl	3-pentyl
1657	1-tetrazolylmethyl	3-pentyl
1658	2-pyridylmethyl	3-pentyl
1659	3-pyridylmethyl	3-pentyl
1660	4-pyridylmethyl	3-pentyl
1661	cyclohexylmethyl	3-pentyl
1662	2-naphthylmethyl	3-pentyl
1663	3-naphthylmethyl	3-pentyl
1664	2-thiophenylmethyl	3-pentyl
1665	4-(1-methyl)piperidinyl-methyl	3-pentyl

Table 4

1666	"(3,4-methylenedioxyphenyl)methyl"	3-pentyl
1667	2-thienylmethyl	3-pentyl
1668	4-biphenylmethyl	3-pentyl
1669	pyrimidinylmethyl	3-pentyl
1670	2-benzothiazolylmethyl	3-pentyl
1671	2-benzothiophenylmethyl	3-pentyl
1672	2-thiomethylethyl	3-pentyl
1673	2-thiomethylmethyl	3-pentyl
1674	2-methylpropyl	3-pentyl
1675	2-methylbutyl	3-pentyl
1676	3-methylbutyl	3-pentyl
1677	cyclopropylmethyl	3-pentyl
1678	cyclobutylmethyl	3-pentyl
1679	cyclopentylmethyl	3-pentyl
1680	p-hydroxyphenylmethyl	3-pentyl
1681	p-nitrophenylmethyl	3-pentyl
1682	p-aminophenylmethyl	3-pentyl
1683	"4-(N,N-dimethylamino)phenylmethyl"	3-pentyl
1684	benzyl	3-methylbutyl
1685	1-pyrollylmethyl	3-methylbutyl
1686	1-pyrazolylmethyl	3-methylbutyl
1687	1-imidazolylmethyl	3-methylbutyl
1688	1-indolylmethyl	3-methylbutyl
1689	1-triazolylmethyl	3-methylbutyl
1690	1-tetrazolylmethyl	3-methylbutyl
1691	2-pyridylmethyl	3-methylbutyl
1692	3-pyridylmethyl	3-methylbutyl
1693	4-pyridylmethyl	3-methylbutyl
1694	cyclohexylmethyl	3-methylbutyl
1695	2-naphthylmethyl	3-methylbutyl
1696	3-naphthylmethyl	3-methylbutyl
1697	2-thiophenylmethyl	3-methylbutyl
1698	4-(1-methyl)piperidinylmethyl	3-methylbutyl
1699	"(3,4-methylenedioxyphenyl)methyl"	3-methylbutyl
1700	2-thienylmethyl	3-methylbutyl
1701	4-biphenylmethyl	3-methylbutyl
1702	pyrimidinylmethyl	3-methylbutyl
1703	2-benzothiazolylmethyl	3-methylbutyl
1704	2-benzothiophenylmethyl	3-methylbutyl
1705	2-thiomethylethyl	3-methylbutyl
1706	2-thiomethylmethyl	3-methylbutyl
1707	2-methylpropyl	3-methylbutyl
1708	2-methylbutyl	3-methylbutyl
1709	3-methylbutyl	3-methylbutyl
1710	cyclopropylmethyl	3-methylbutyl
1711	cyclobutylmethyl	3-methylbutyl
1712	cyclopentylmethyl	3-methylbutyl
1713	p-hydroxyphenylmethyl	3-methylbutyl
1714	p-nitrophenylmethyl	3-methylbutyl
1715	p-aminophenylmethyl	3-methylbutyl
1716	"4-(N,N-dimethylamino)phenylmethyl"	3-methylbutyl
1717	benzyl	2-methylbutyl

Table 4

1718	1-pyrolylmethyl	2-methylbutyl
1719	1-pyrazolylmethyl	2-methylbutyl
1720	1-imidazolylmethyl	2-methylbutyl
1721	1-indolylmethyl	2-methylbutyl
1722	1-triazolylmethyl	2-methylbutyl
1723	1-tetrazolylmethyl	2-methylbutyl
1724	2-pyridylmethyl	2-methylbutyl
1725	3-pyridylmethyl	2-methylbutyl
1726	4-pyridylmethyl	2-methylbutyl
1727	cyclohexylmethyl	2-methylbutyl
1728	2-naphthylmethyl	2-methylbutyl
1729	3-naphthylmethyl	2-methylbutyl
1730	2-thiophenylmethyl	2-methylbutyl
1731	4-(1-methyl)piperidinyl- methyl	2-methylbutyl
1732	"(3,4- methylenedioxyphenyl)methyl"	2-methylbutyl
1733	2-thienylmethyl	2-methylbutyl
1734	4-biphenylmethyl	2-methylbutyl
1735	pyrimidinylmethyl	2-methylbutyl
1736	2-benzothiazolylmethyl	2-methylbutyl
1737	2-benzothiophenylmethyl	2-methylbutyl
1738	2-thiomethylethyl	2-methylbutyl
1739	2-thiomethylmethyl	2-methylbutyl
1740	2-methylpropyl	2-methylbutyl
1741	2-methylbutyl	2-methylbutyl
1742	3-methylbutyl	2-methylbutyl
1743	cyclopropylmethyl	2-methylbutyl
1744	cyclobutylmethyl	2-methylbutyl
1745	cyclopentylmethyl	2-methylbutyl
1746	p-hydroxyphenylmethyl	2-methylbutyl
1747	p-nitrophenylmethyl	2-methylbutyl
1748	p-aminophenylmethyl	2-methylbutyl
1749	"4-(N,N- dimethylamino)phenylmethyl" benzyl	propargyl
1750	1-pyrolylmethyl	propargyl
1751	1-pyrazolylmethyl	propargyl
1752	1-imidazolylmethyl	propargyl
1753	1-indolylmethyl	propargyl
1754	1-triazolylmethyl	propargyl
1755	1-tetrazolylmethyl	propargyl
1756	2-pyridylmethyl	propargyl
1757	3-pyridylmethyl	propargyl
1758	4-pyridylmethyl	propargyl
1759	cyclohexylmethyl	propargyl
1760	2-naphthylmethyl	propargyl
1761	3-naphthylmethyl	propargyl
1762	2-thiophenylmethyl	propargyl
1763	4-(1-methyl)piperidinyl- methyl	propargyl
1764	"(3,4- methylenedioxyphenyl)methyl"	propargyl
1765	2-thienylmethyl	propargyl
1766	4-biphenylmethyl	propargyl
1767	pyrimidinylmethyl	propargyl
1768	2-benzothiazolylmethyl	propargyl
1769		

Table 4

1770	2-benzothiophenylmethyl	propargyl
1771	2-thiomethylethyl	propargyl
1772	2-thiomethylmethyl	propargyl
1773	2-methylpropyl	propargyl
1774	2-methylbutyl	propargyl
1775	3-methylbutyl	propargyl
1776	cyclopropylmethyl	propargyl
1777	cyclobutylmethyl	propargyl
1778	cyclopentylmethyl	propargyl
1779	p-hydroxyphenylmethyl	propargyl
1780	p-nitrophenylmethyl	propargyl
1781	p-aminophenylmethyl	propargyl
1782	"4-(N,N-dimethylamino)phenylmethyl"	propargyl
1783	benzyl	cyclobutyl
1784	1-pyrolylmethyl	cyclobutyl
1785	1-pyrazolylmethyl	cyclobutyl
1786	1-imidazolylmethyl	cyclobutyl
1787	1-indolylmethyl	cyclobutyl
1788	1-triazolylmethyl	cyclobutyl
1789	1-tetrazolylmethyl	cyclobutyl
1790	2-pyridylmethyl	cyclobutyl
1791	3-pyridylmethyl	cyclobutyl
1792	4-pyridylmethyl	cyclobutyl
1793	cyclohexylmethyl	cyclobutyl
1794	2-naphthylmethyl	cyclobutyl
1795	3-naphthylmethyl	cyclobutyl
1796	2-thiophenylmethyl	cyclobutyl
1797	4-(1-methyl)piperidinyl-methyl	cyclobutyl
1798	"(3,4-methylenedioxyphenyl)methyl"	cyclobutyl
1799	2-thienylmethyl	cyclobutyl
1800	4-biphenylmethyl	cyclobutyl
1801	pyrimidinylmethyl	cyclobutyl
1802	2-benzothiazolylmethyl	cyclobutyl
1803	2-benzothiophenylmethyl	cyclobutyl
1804	2-thiomethylethyl	cyclobutyl
1805	2-thiomethylmethyl	cyclobutyl
1806	2-methylpropyl	cyclobutyl
1807	2-methylbutyl	cyclobutyl
1808	3-methylbutyl	cyclobutyl
1809	cyclopropylmethyl	cyclobutyl
1810	cyclobutylmethyl	cyclobutyl
1811	cyclopentylmethyl	cyclobutyl
1812	p-hydroxyphenylmethyl	cyclobutyl
1813	p-nitrophenylmethyl	cyclobutyl
1814	p-aminophenylmethyl	cyclobutyl
1815	"4-(N,N-dimethylamino)phenylmethyl"	cyclobutyl
1816	benzyl	cyclobutylmethyl
1817	1-pyrolylmethyl	cyclobutylmethyl
1818	1-pyrazolylmethyl	cyclobutylmethyl
1819	1-imidazolylmethyl	cyclobutylmethyl
1820	1-indolylmethyl	cyclobutylmethyl
1821	1-triazolylmethyl	cyclobutylmethyl
1822	1-tetrazolylmethyl	cyclobutylmethyl

Table 4

1823	2-pyridylmethyl	cyclobutylmethyl
1824	3-pyridylmethyl	cyclobutylmethyl
1825	4-pyridylmethyl	cyclobutylmethyl
1826	cyclohexylmethyl	cyclobutylmethyl
1827	2-naphthylmethyl	cyclobutylmethyl
1828	3-naphthylmethyl	cyclobutylmethyl
1829	2-thiophenylmethyl	cyclobutylmethyl
1830	4-(1-methyl)piperidinyl-	cyclobutylmethyl
	methyl	
1831	"(3,4-	cyclobutylmethyl
	methylenedioxyphenyl)methyl"	
1832	2-thienylmethyl	cyclobutylmethyl
1833	4-biphenylmethyl	cyclobutylmethyl
1834	pyrimidinylmethyl	cyclobutylmethyl
1835	2-benzothiazolylmethyl	cyclobutylmethyl
1836	2-benzothiophenylmethyl	cyclobutylmethyl
1837	2-thiomethylethyl	cyclobutylmethyl
1838	2-thiomethylmethyl	cyclobutylmethyl
1839	2-methylpropyl	cyclobutylmethyl
1840	2-methylbutyl	cyclobutylmethyl
1841	3-methylbutyl	cyclobutylmethyl
1842	cyclopropylmethyl	cyclobutylmethyl
1843	cyclobutylmethyl	cyclobutylmethyl
1844	cyclopentylmethyl	cyclobutylmethyl
1845	p-hydroxyphenylmethyl	cyclobutylmethyl
1846	p-nitrophenylmethyl	cyclobutylmethyl
1847	p-aminophenylmethyl	cyclobutylmethyl
1848	"4-(N,N-	cyclobutylmethyl
	dimethylamino)phenylmethyl"	
1849	benzyl	cyclopentyl
1850	1-pyrollylmethyl	cyclopentyl
1851	1-pyrazolylmethyl	cyclopentyl
1852	1-imidazolylmethyl	cyclopentyl
1853	1-indolylmethyl	cyclopentyl
1854	1-triazolylmethyl	cyclopentyl
1855	1-tetrazolylmethyl	cyclopentyl
1856	2-pyridylmethyl	cyclopentyl
1857	3-pyridylmethyl	cyclopentyl
1858	4-pyridylmethyl	cyclopentyl
1859	cyclohexylmethyl	cyclopentyl
1860	2-naphthylmethyl	cyclopentyl
1861	3-naphthylmethyl	cyclopentyl
1862	2-thiophenylmethyl	cyclopentyl
1863	4-(1-methyl)piperidinyl-	cyclopentyl
	methyl	
1864	"(3,4-	cyclopentyl
	methylenedioxyphenyl)methyl"	
1865	2-thienylmethyl	cyclopentyl
1866	4-biphenylmethyl	cyclopentyl
1867	pyrimidinylmethyl	cyclopentyl
1868	2-benzothiazolylmethyl	cyclopentyl
1869	2-benzothiophenylmethyl	cyclopentyl
1870	2-thiomethylethyl	cyclopentyl
1871	2-thiomethylmethyl	cyclopentyl
1872	2-methylpropyl	cyclopentyl
1873	2-methylbutyl	cyclopentyl
1874	3-methylbutyl	cyclopentyl

Table 4

1875	cyclopropylmethyl	cyclopentyl
1876	cyclobutylmethyl	cyclopentyl
1877	cyclopentylmethyl	cyclopentyl
1878	p-hydroxyphenylmethyl	cyclopentyl
1879	p-nitrophenylmethyl	cyclopentyl
1880	p-aminophenylmethyl	cyclopentyl
1881	"4-(N,N-dimethylamino)phenylmethyl"	cyclopentyl
1882	benzyl	cyclopentylmethyl
1883	1-pyrolylmethyl	cyclopentylmethyl
1884	1-pyrazolylmethyl	cyclopentylmethyl
1885	1-imidazolylmethyl	cyclopentylmethyl
1886	1-indolylmethyl	cyclopentylmethyl
1887	1-triazolylmethyl	cyclopentylmethyl
1888	1-tetrazolylmethyl	cyclopentylmethyl
1889	2-pyridylmethyl	cyclopentylmethyl
1890	3-pyridylmethyl	cyclopentylmethyl
1891	4-pyridylmethyl	cyclopentylmethyl
1892	cyclohexylmethyl	cyclopentylmethyl
1893	2-naphthylmethyl	cyclopentylmethyl
1894	3-naphthylmethyl	cyclopentylmethyl
1895	2-thiophenylmethyl	cyclopentylmethyl
1896	4-(1-methyl)piperidinyl-methyl	cyclopentylmethyl
1897	"(3,4-methylenedioxyphenyl)methyl"	cyclopentylmethyl
1898	2-thienylmethyl	cyclopentylmethyl
1899	4-biphenylmethyl	cyclopentylmethyl
1900	pyrimidinylmethyl	cyclopentylmethyl
1901	2-benzothiazolylmethyl	cyclopentylmethyl
1902	2-benzothiophenylmethyl	cyclopentylmethyl
1903	2-thiomethylethyl	cyclopentylmethyl
1904	2-thiomethylmethyl	cyclopentylmethyl
1905	2-methylpropyl	cyclopentylmethyl
1906	2-methylbutyl	cyclopentylmethyl
1907	3-methylbutyl	cyclopentylmethyl
1908	cyclopropylmethyl	cyclopentylmethyl
1909	cyclobutylmethyl	cyclopentylmethyl
1910	cyclopentylmethyl	cyclopentylmethyl
1911	p-hydroxyphenylmethyl	cyclopentylmethyl
1912	p-nitrophenylmethyl	cyclopentylmethyl
1913	p-aminophenylmethyl	cyclopentylmethyl
1914	"4-(N,N-dimethylamino)phenylmethyl"	cyclopentylmethyl
1915	benzyl	1-hexyl
1916	1-pyrolylmethyl	1-hexyl
1917	1-pyrazolylmethyl	1-hexyl
1918	1-imidazolylmethyl	1-hexyl
1919	1-indolylmethyl	1-hexyl
1920	1-triazolylmethyl	1-hexyl
1921	1-tetrazolylmethyl	1-hexyl
1922	2-pyridylmethyl	1-hexyl
1923	3-pyridylmethyl	1-hexyl
1924	4-pyridylmethyl	1-hexyl
1925	cyclohexylmethyl	1-hexyl
1926	2-naphthylmethyl	1-hexyl
1927	3-naphthylmethyl	1-hexyl

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Table 4

1928	2-thiophenylmethyl	1-hexyl
1929	4-(1-methyl)piperidinyl-methyl	1-hexyl
1930	"(3,4-methylenedioxyphenyl)methyl"	1-hexyl
1931	2-thienylmethyl	1-hexyl
1932	4-biphenylmethyl	1-hexyl
1933	pyrimidinylmethyl	1-hexyl
1934	2-benzothiazolylmethyl	1-hexyl
1935	2-benzothiophenylmethyl	1-hexyl
1936	2-thiomethylethyl	1-hexyl
1937	2-thiomethylmethyl	1-hexyl
1938	2-methylpropyl	1-hexyl
1939	2-methylbutyl	1-hexyl
1940	3-methylbutyl	1-hexyl
1941	cyclopropylmethyl	1-hexyl
1942	cyclobutylmethyl	1-hexyl
1943	cyclopentylmethyl	1-hexyl
1944	p-hydroxyphenylmethyl	1-hexyl
1945	p-nitrophenylmethyl	1-hexyl
1946	p-aminophenylmethyl	1-hexyl
1947	"4-(N,N-dimethylamino)phenylmethyl" benzyl	4-methylpentyl
1948	1-pyrollylmethyl	4-methylpentyl
1949	1-pyrazolylmethyl	4-methylpentyl
1950	1-imidazolylmethyl	4-methylpentyl
1951	1-indolylmethyl	4-methylpentyl
1952	1-triazolylmethyl	4-methylpentyl
1953	1-tetrazolylmethyl	4-methylpentyl
1954	2-pyridylmethyl	4-methylpentyl
1955	3-pyridylmethyl	4-methylpentyl
1956	4-pyridylmethyl	4-methylpentyl
1957	cyclohexylmethyl	4-methylpentyl
1958	2-naphthylmethyl	4-methylpentyl
1959	3-naphthylmethyl	4-methylpentyl
1960	2-thiophenylmethyl	4-methylpentyl
1961	4-(1-methyl)piperidinyl-methyl	4-methylpentyl
1962	"(3,4-methylenedioxyphenyl)methyl"	4-methylpentyl
1963	2-thienylmethyl	4-methylpentyl
1964	4-biphenylmethyl	4-methylpentyl
1965	pyrimidinylmethyl	4-methylpentyl
1966	2-benzothiazolylmethyl	4-methylpentyl
1967	2-benzothiophenylmethyl	4-methylpentyl
1968	2-thiomethylethyl	4-methylpentyl
1969	2-thiomethylmethyl	4-methylpentyl
1970	2-methylpropyl	4-methylpentyl
1971	2-methylbutyl	4-methylpentyl
1972	3-methylbutyl	4-methylpentyl
1973	cyclopropylmethyl	4-methylpentyl
1974	cyclobutylmethyl	4-methylpentyl
1975	cyclopentylmethyl	4-methylpentyl
1976	p-hydroxyphenylmethyl	4-methylpentyl
1977	p-nitrophenylmethyl	4-methylpentyl
1978	p-aminophenylmethyl	4-methylpentyl
1979		

Table 4

1980	"4-(N,N-dimethylamino)phenylmethyl"	4-methylpentyl
1981	benzyl	3-methylpentyl
1982	1-pyrolylmethyl	3-methylpentyl
1983	1-pyrazolylmethyl	3-methylpentyl
1984	1-imidazolylmethyl	3-methylpentyl
1985	1-indolylmethyl	3-methylpentyl
1986	1-triazolylmethyl	3-methylpentyl
1987	1-tetrazolylmethyl	3-methylpentyl
1988	2-pyridylmethyl	3-methylpentyl
1989	3-pyridylmethyl	3-methylpentyl
1990	4-pyridylmethyl	3-methylpentyl
1991	cyclohexylmethyl	3-methylpentyl
1992	2-naphthylmethyl	3-methylpentyl
1993	3-naphthylmethyl	3-methylpentyl
1994	2-thiophenylmethyl	3-methylpentyl
1995	4-(1-methyl)piperidinyl-methyl	3-methylpentyl
1996	"(3,4-methylenedioxyphenyl)methyl"	3-methylpentyl
1997	2-thienylmethyl	3-methylpentyl
1998	4-biphenylmethyl	3-methylpentyl
1999	pyrimidinylmethyl	3-methylpentyl
2000	2-benzothiazolylmethyl	3-methylpentyl
2001	2-benzothiophenylmethyl	3-methylpentyl
2002	2-thiomethylethyl	3-methylpentyl
2003	2-thiomethylmethyl	3-methylpentyl
2004	2-methylpropyl	3-methylpentyl
2005	2-methylbutyl	3-methylpentyl
2006	3-methylbutyl	3-methylpentyl
2007	cyclopropylmethyl	3-methylpentyl
2008	cyclobutylmethyl	3-methylpentyl
2009	cyclopentylmethyl	3-methylpentyl
2010	p-hydroxyphenylmethyl	3-methylpentyl
2011	p-nitrophenylmethyl	3-methylpentyl
2012	p-aminophenylmethyl	3-methylpentyl
2013	"4-(N,N-dimethylamino)phenylmethyl"	isopropyl
2014	benzyl	isopropyl
2015	1-pyrolylmethyl	isopropyl
2016	1-pyrazolylmethyl	isopropyl
2017	1-imidazolylmethyl	isopropyl
2018	1-indolylmethyl	isopropyl
2019	1-triazolylmethyl	isopropyl
2020	1-tetrazolylmethyl	isopropyl
2021	2-pyridylmethyl	isopropyl
2022	3-pyridylmethyl	isopropyl
2023	4-pyridylmethyl	isopropyl
2024	cyclohexylmethyl	isopropyl
2025	2-naphthylmethyl	isopropyl
2026	3-naphthylmethyl	isopropyl
2027	2-thiophenylmethyl	isopropyl
2028	4-(1-methyl)piperidinyl-methyl	isopropyl
2029	"(3,4-methylenedioxyphenyl)methyl"	isopropyl
2030	2-thienylmethyl	isopropyl

Table 4

2031	4-biphenylmethyl	isopropyl
2032	pyrimidinylmethyl	isopropyl
2033	2-benzothiazolylmethyl	isopropyl
2034	2-benzothiophenylmethyl	isopropyl
2035	2-thiomethylethyl	isopropyl
2036	2-thiomethylmethyl	isopropyl
2037	2-methylpropyl	isopropyl
2038	2-methylbutyl	isopropyl
2039	3-methylbutyl	isopropyl
2040	cyclopropylmethyl	isopropyl
2041	cyclobutylmethyl	isopropyl
2042	cyclopentylmethyl	isopropyl
2043	p-hydroxyphenylmethyl	isopropyl
2044	p-nitrophenylmethyl	isopropyl
2045	p-aminophenylmethyl	isopropyl
2046	"4-(N,N-dimethylamino)phenylmethyl"	isopropyl
2047	benzyl	3-methylbutyl
2048	1-pyrolylmethyl	3-methylbutyl
2049	1-pyrazolylmethyl	3-methylbutyl
2050	1-imidazolylmethyl	3-methylbutyl
2051	1-indolylmethyl	3-methylbutyl
2052	1-triazolylmethyl	3-methylbutyl
2053	1-tetrazolylmethyl	3-methylbutyl
2054	2-pyridylmethyl	3-methylbutyl
2055	3-pyridylmethyl	3-methylbutyl
2056	4-pyridylmethyl	3-methylbutyl
2057	cyclohexylmethyl	3-methylbutyl
2058	2-naphthylmethyl	3-methylbutyl
2059	3-naphthylmethyl	3-methylbutyl
2060	2-thiophenylmethyl	3-methylbutyl
2061	4-(1-methyl)piperidinyl-methyl	3-methylbutyl
2062	"(3,4-methylenedioxyphenyl)methyl"	3-methylbutyl
2063	2-thienylmethyl	3-methylbutyl
2064	4-biphenylmethyl	3-methylbutyl
2065	pyrimidinylmethyl	3-methylbutyl
2066	2-benzothiazolylmethyl	3-methylbutyl
2067	2-benzothiophenylmethyl	3-methylbutyl
2068	2-thiomethylethyl	3-methylbutyl
2069	2-thiomethylmethyl	3-methylbutyl
2070	2-methylpropyl	3-methylbutyl
2071	2-methylbutyl	3-methylbutyl
2072	3-methylbutyl	3-methylbutyl
2073	cyclopropylmethyl	3-methylbutyl
2074	cyclobutylmethyl	3-methylbutyl
2075	cyclopentylmethyl	3-methylbutyl
2076	p-hydroxyphenylmethyl	3-methylbutyl
2077	p-nitrophenylmethyl	3-methylbutyl
2078	p-aminophenylmethyl	3-methylbutyl
2079	"4-(N,N-dimethylamino)phenylmethyl"	3-methylbutyl
2080	benzyl	2-phenylethyl
2081	1-pyrolylmethyl	2-phenylethyl
2082	1-pyrazolylmethyl	2-phenylethyl
2083	1-imidazolylmethyl	2-phenylethyl

Table 4

2084	1-indolylmethyl	2-phenylethyl
2085	1-triazolylmethyl	2-phenylethyl
2086	1-tetrazolylmethyl	2-phenylethyl
2087	2-pyridylmethyl	2-phenylethyl
2088	3-pyridylmethyl	2-phenylethyl
2089	4-pyridylmethyl	2-phenylethyl
2090	cyclohexylmethyl	2-phenylethyl
2091	2-naphthylmethyl	2-phenylethyl
2092	3-naphthylmethyl	2-phenylethyl
2093	2-thiophenylmethyl	2-phenylethyl
2094	4-(1-methyl)piperidinyl- methyl	2-phenylethyl
2095	"(3,4- methylenedioxyphenyl)methyl"	2-phenylethyl
2096	2-thienylmethyl	2-phenylethyl
2097	4-biphenylmethyl	2-phenylethyl
2098	pyrimidinylmethyl	2-phenylethyl
2099	2-benzothiazolylmethyl	2-phenylethyl
2100	2-benzothiophenylmethyl	2-phenylethyl
2101	2-thiomethylethyl	2-phenylethyl
2102	2-thiomethylmethyl	2-phenylethyl
2103	2-methylpropyl	2-phenylethyl
2104	2-methylbutyl	2-phenylethyl
2105	3-methylbutyl	2-phenylethyl
2106	cyclopropylmethyl	2-phenylethyl
2107	cyclobutylmethyl	2-phenylethyl
2108	cyclopentylmethyl	2-phenylethyl
2109	p-hydroxyphenylmethyl	2-phenylethyl
2110	p-nitrophenylmethyl	2-phenylethyl
2111	p-aminophenylmethyl	2-phenylethyl
2112	"4-(N,N- dimethylamino)phenylmethyl" benzyl	2-phenylethyl
2113	1-pyrollylmethyl	3-phenylpropyl
2114	1-pyrazolylmethyl	3-phenylpropyl
2115	1-imidazolylmethyl	3-phenylpropyl
2116	1-indolylmethyl	3-phenylpropyl
2117	1-triazolylmethyl	3-phenylpropyl
2118	1-tetrazolylmethyl	3-phenylpropyl
2119	2-pyridylmethyl	3-phenylpropyl
2120	3-pyridylmethyl	3-phenylpropyl
2121	4-pyridylmethyl	3-phenylpropyl
2122	cyclohexylmethyl	3-phenylpropyl
2123	2-naphthylmethyl	3-phenylpropyl
2124	3-naphthylmethyl	3-phenylpropyl
2125	2-thiophenylmethyl	3-phenylpropyl
2126	4-(1-methyl)piperidinyl- methyl	3-phenylpropyl
2127	"(3,4- methylenedioxyphenyl)methyl"	3-phenylpropyl
2128	2-thienylmethyl	3-phenylpropyl
2129	4-biphenylmethyl	3-phenylpropyl
2130	pyrimidinylmethyl	3-phenylpropyl
2131	2-benzothiazolylmethyl	3-phenylpropyl
2132	2-benzothiophenylmethyl	3-phenylpropyl
2133	2-thiomethylethyl	3-phenylpropyl
2134	2-thiomethylmethyl	3-phenylpropyl
2135		

Table 4

2136	2-methylpropyl	3-phenylpropyl
2137	2-methylbutyl	3-phenylpropyl
2138	3-methylbutyl	3-phenylpropyl
2139	cyclopropylmethyl	3-phenylpropyl
2140	cyclobutylmethyl	3-phenylpropyl
2141	cyclopentylmethyl	3-phenylpropyl
2142	p-hydroxyphenylmethyl	3-phenylpropyl
2143	p-nitrophenylmethyl	3-phenylpropyl
2144	p-aminophenylmethyl	3-phenylpropyl
2145	"4-(N,N-dimethylamino)phenylmethyl"	3-phenylpropyl
2146	benzyl	"2-(N,N-dimethylamino)ethyl"
2147	1-pyrrolylmethyl	"2-(N,N-dimethylamino)ethyl"
2148	1-pyrazolylmethyl	"2-(N,N-dimethylamino)ethyl"
2149	1-imidazolylmethyl	"2-(N,N-dimethylamino)ethyl"
2150	1-indolylmethyl	"2-(N,N-dimethylamino)ethyl"
2151	1-triazolylmethyl	"2-(N,N-dimethylamino)ethyl"
2152	1-tetrazolylmethyl	"2-(N,N-dimethylamino)ethyl"
2153	2-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
2154	3-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
2155	4-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
2156	cyclohexylmethyl	"2-(N,N-dimethylamino)ethyl"
2157	2-naphthylmethyl	"2-(N,N-dimethylamino)ethyl"
2158	3-naphthylmethyl	"2-(N,N-dimethylamino)ethyl"
2159	2-thiophenylmethyl	"2-(N,N-dimethylamino)ethyl"
2160	4-(1-methyl)piperidinyl-methyl	"2-(N,N-dimethylamino)ethyl"
2161	"(3,4-methylenedioxyphenyl)methyl"	"2-(N,N-dimethylamino)ethyl"
2162	2-thienylmethyl	"2-(N,N-dimethylamino)ethyl"
2163	4-biphenylmethyl	"2-(N,N-dimethylamino)ethyl"
2164	pyrimidinylmethyl	"2-(N,N-dimethylamino)ethyl"
2165	2-benzothiazolylmethyl	"2-(N,N-dimethylamino)ethyl"
2166	2-benzothiophenylmethyl	"2-(N,N-dimethylamino)ethyl"
2167	2-thiomethylethyl	"2-(N,N-dimethylamino)ethyl"
2168	2-thiomethylmethyl	"2-(N,N-dimethylamino)ethyl"
2169	2-methylpropyl	"2-(N,N-dimethylamino)ethyl"
2170	2-methylbutyl	"2-(N,N-dimethylamino)ethyl"
2171	3-methylbutyl	"2-(N,N-dimethylamino)ethyl"
2172	cyclopropylmethyl	"2-(N,N-dimethylamino)ethyl"
2173	cyclobutylmethyl	"2-(N,N-dimethylamino)ethyl"
2174	cyclopentylmethyl	"2-(N,N-dimethylamino)ethyl"
2175	p-hydroxyphenylmethyl	"2-(N,N-dimethylamino)ethyl"
2176	p-nitrophenylmethyl	"2-(N,N-dimethylamino)ethyl"
2177	p-aminophenylmethyl	"2-(N,N-dimethylamino)ethyl"
2178	"4-(N,N-dimethylamino)phenylmethyl"	"2-(N,N-dimethylamino)ethyl"
2179	benzyl	3-oxetanylmethyl
2180	1-pyrrolylmethyl	3-oxetanylmethyl
2181	1-pyrazolylmethyl	3-oxetanylmethyl
2182	1-imidazolylmethyl	3-oxetanylmethyl
2183	1-indolylmethyl	3-oxetanylmethyl
2184	1-triazolylmethyl	3-oxetanylmethyl
2185	1-tetrazolylmethyl	3-oxetanylmethyl
2186	2-pyridylmethyl	3-oxetanylmethyl
2187	3-pyridylmethyl	3-oxetanylmethyl
2188	4-pyridylmethyl	3-oxetanylmethyl

Table 4

2189	cyclohexylmethyl	3-oxetanylmethyl
2190	2-naphthylmethyl	3-oxetanylmethyl
2191	3-naphthylmethyl	3-oxetanylmethyl
2192	2-thiophenylmethyl	3-oxetanylmethyl
2193	4-(1-methyl)piperidinyl-	3-oxetanylmethyl
	methyl	
2194	"(3,4-	3-oxetanylmethyl
	methylenedioxyphenyl)methyl"	
2195	2-thienylmethyl	3-oxetanylmethyl
2196	4-biphenylmethyl	3-oxetanylmethyl
2197	pyrimidinylmethyl	3-oxetanylmethyl
2198	2-benzothiazolylmethyl	3-oxetanylmethyl
2199	2-benzothiophenylmethyl	3-oxetanylmethyl
2200	2-thiomethylethyl	3-oxetanylmethyl
2201	2-thiomethylmethyl	3-oxetanylmethyl
2202	2-methylpropyl	3-oxetanylmethyl
2203	2-methylbutyl	3-oxetanylmethyl
2204	3-methylbutyl	3-oxetanylmethyl
2205	cyclopropylmethyl	3-oxetanylmethyl
2206	cyclobutylmethyl	3-oxetanylmethyl
2207	cyclopentylmethyl	3-oxetanylmethyl
2208	p-hydroxyphenylmethyl	3-oxetanylmethyl
2209	p-nitrophenylmethyl	3-oxetanylmethyl
2210	p-aminophenylmethyl	3-oxetanylmethyl
2211	"4-(N,N-	3-oxetanylmethyl
	dimethylamino)phenylmethyl"	
2212	benzyl	2-tetrahydrofuranyl
2213	1-pyrrolylmethyl	2-tetrahydrofuranyl
2214	1-pyrazolylmethyl	2-tetrahydrofuranyl
2215	1-imidazolylmethyl	2-tetrahydrofuranyl
2216	1-indolylmethyl	2-tetrahydrofuranyl
2217	1-triazolylmethyl	2-tetrahydrofuranyl
2218	1-tetrazolylmethyl	2-tetrahydrofuranyl
2219	2-pyridylmethyl	2-tetrahydrofuranyl
2220	3-pyridylmethyl	2-tetrahydrofuranyl
2221	4-pyridylmethyl	2-tetrahydrofuranyl
2222	cyclohexylmethyl	2-tetrahydrofuranyl
2223	2-naphthylmethyl	2-tetrahydrofuranyl
2224	3-naphthylmethyl	2-tetrahydrofuranyl
2225	2-thiophenylmethyl	2-tetrahydrofuranyl
2226	4-(1-methyl)piperidinyl-	2-tetrahydrofuranyl
	methyl	
2227	"(3,4-	2-tetrahydrofuranyl
	methylenedioxyphenyl)methyl"	
2228	2-thienylmethyl	2-tetrahydrofuranyl
2229	4-biphenylmethyl	2-tetrahydrofuranyl
2230	pyrimidinylmethyl	2-tetrahydrofuranyl
2231	2-benzothiazolylmethyl	2-tetrahydrofuranyl
2232	2-benzothiophenylmethyl	2-tetrahydrofuranyl
2233	2-thiomethylethyl	2-tetrahydrofuranyl
2234	2-thiomethylmethyl	2-tetrahydrofuranyl
2235	2-methylpropyl	2-tetrahydrofuranyl
2236	2-methylbutyl	2-tetrahydrofuranyl
2237	3-methylbutyl	2-tetrahydrofuranyl
2238	cyclopropylmethyl	2-tetrahydrofuranyl
2239	cyclobutylmethyl	2-tetrahydrofuranyl
2240	cyclopentylmethyl	2-tetrahydrofuranyl

Table 4

2241	p-hydroxyphenylmethyl	2-tetrahydrofuranyl
2242	p-nitrophenylmethyl	2-tetrahydrofuranyl
2243	p-aminophenylmethyl	2-tetrahydrofuranyl
2244	"4-(N,N-dimethylamino)phenylmethyl"	2-tetrahydrofuranyl
2245	benzyl	2-methoxypropyl
2246	1-pyrolylmethyl	2-methoxypropyl
2247	1-pyrazolylmethyl	2-methoxypropyl
2248	1-imidazolylmethyl	2-methoxypropyl
2249	1-indolylmethyl	2-methoxypropyl
2250	1-triazolylmethyl	2-methoxypropyl
2251	1-tetrazolylmethyl	2-methoxypropyl
2252	2-pyridylmethyl	2-methoxypropyl
2253	3-pyridylmethyl	2-methoxypropyl
2254	4-pyridylmethyl	2-methoxypropyl
2255	cyclohexylmethyl	2-methoxypropyl
2256	2-naphthylmethyl	2-methoxypropyl
2257	3-naphthylmethyl	2-methoxypropyl
2258	2-thiophenylmethyl	2-methoxypropyl
2259	4-(1-methyl)piperidinyl-methyl	2-methoxypropyl
2260	"(3,4-methylenedioxyphenyl)methyl"	2-methoxypropyl
2261	2-thienylmethyl	2-methoxypropyl
2262	4-biphenylmethyl	2-methoxypropyl
2263	pyrimidinylmethyl	2-methoxypropyl
2264	2-benzothiazolylmethyl	2-methoxypropyl
2265	2-benzothiophenylmethyl	2-methoxypropyl
2266	2-thiomethylethyl	2-methoxypropyl
2267	2-thiomethylmethyl	2-methoxypropyl
2268	2-methylpropyl	2-methoxypropyl
2269	2-methylbutyl	2-methoxypropyl
2270	3-methylbutyl	2-methoxypropyl
2271	cyclopropylmethyl	2-methoxypropyl
2272	cyclobutylmethyl	2-methoxypropyl
2273	cyclopentylmethyl	2-methoxypropyl
2274	p-hydroxyphenylmethyl	2-methoxypropyl
2275	p-nitrophenylmethyl	2-methoxypropyl
2276	p-aminophenylmethyl	2-methoxypropyl
2277	"4-(N,N-dimethylamino)phenylmethyl"	2-methoxypropyl
2278	benzyl	2-ethoxyethyl
2279	1-pyrolylmethyl	2-ethoxyethyl
2280	1-pyrazolylmethyl	2-ethoxyethyl
2281	1-imidazolylmethyl	2-ethoxyethyl
2282	1-indolylmethyl	2-ethoxyethyl
2283	1-triazolylmethyl	2-ethoxyethyl
2284	1-tetrazolylmethyl	2-ethoxyethyl
2285	2-pyridylmethyl	2-ethoxyethyl
2286	3-pyridylmethyl	2-ethoxyethyl
2287	4-pyridylmethyl	2-ethoxyethyl
2288	cyclohexylmethyl	2-ethoxyethyl
2289	2-naphthylmethyl	2-ethoxyethyl
2290	3-naphthylmethyl	2-ethoxyethyl
2291	2-thiophenylmethyl	2-ethoxyethyl
2292	4-(1-methyl)piperidinyl-methyl	2-ethoxyethyl

Table 4

2293	"(3,4-methylenedioxyphenyl)methyl"	2-ethoxyethyl
2294	2-thienylmethyl	2-ethoxyethyl
2295	4-biphenylmethyl	2-ethoxyethyl
2296	pyrimidinylmethyl	2-ethoxyethyl
2297	2-benzothiazolylmethyl	2-ethoxyethyl
2298	2-benzothiophenylmethyl	2-ethoxyethyl
2299	2-thiomethylethyl	2-ethoxyethyl
2300	2-thiomethylmethyl	2-ethoxyethyl
2301	2-methylpropyl	2-ethoxyethyl
2302	2-methylbutyl	2-ethoxyethyl
2303	3-methylbutyl	2-ethoxyethyl
2304	cyclopropylmethyl	2-ethoxyethyl
2305	cyclobutylmethyl	2-ethoxyethyl
2306	cyclopentylmethyl	2-ethoxyethyl
2307	p-hydroxyphenylmethyl	2-ethoxyethyl
2308	p-nitrophenylmethyl	2-ethoxyethyl
2309	p-aminophenylmethyl	2-ethoxyethyl
2310	"4-(N,N-dimethylamino)phenylmethyl"	2-ethoxyethyl
2311	benzyl	2-(1-pyrolyl)ethyl
2312	1-pyrollylmethyl	2-(1-pyrolyl)ethyl
2313	1-pyrazolylmethyl	2-(1-pyrolyl)ethyl
2314	1-imidazolylmethyl	2-(1-pyrolyl)ethyl
2315	1-indolylmethyl	2-(1-pyrolyl)ethyl
2316	1-triazolylmethyl	2-(1-pyrolyl)ethyl
2317	1-tetrazolylmethyl	2-(1-pyrolyl)ethyl
2318	2-pyridylmethyl	2-(1-pyrolyl)ethyl
2319	3-pyridylmethyl	2-(1-pyrolyl)ethyl
2320	4-pyridylmethyl	2-(1-pyrolyl)ethyl
2321	cyclohexylmethyl	2-(1-pyrolyl)ethyl
2322	2-naphthylmethyl	2-(1-pyrolyl)ethyl
2323	3-naphthylmethyl	2-(1-pyrolyl)ethyl
2324	2-thiophenylmethyl	2-(1-pyrolyl)ethyl
2325	4-(1-methyl)piperidinylmethyl	2-(1-pyrolyl)ethyl
2326	"(3,4-methylenedioxyphenyl)methyl"	2-(1-pyrolyl)ethyl
2327	2-thienylmethyl	2-(1-pyrolyl)ethyl
2328	4-biphenylmethyl	2-(1-pyrolyl)ethyl
2329	pyrimidinylmethyl	2-(1-pyrolyl)ethyl
2330	2-benzothiazolylmethyl	2-(1-pyrolyl)ethyl
2331	2-benzothiophenylmethyl	2-(1-pyrolyl)ethyl
2332	2-thiomethylethyl	2-(1-pyrolyl)ethyl
2333	2-thiomethylmethyl	2-(1-pyrolyl)ethyl
2334	2-methylpropyl	2-(1-pyrolyl)ethyl
2335	2-methylbutyl	2-(1-pyrolyl)ethyl
2336	3-methylbutyl	2-(1-pyrolyl)ethyl
2337	cyclopropylmethyl	2-(1-pyrolyl)ethyl
2338	cyclobutylmethyl	2-(1-pyrolyl)ethyl
2339	cyclopentylmethyl	2-(1-pyrolyl)ethyl
2340	p-hydroxyphenylmethyl	2-(1-pyrolyl)ethyl
2341	p-nitrophenylmethyl	2-(1-pyrolyl)ethyl
2342	p-aminophenylmethyl	2-(1-pyrolyl)ethyl
2343	"4-(N,N-dimethylamino)phenylmethyl"	2-(1-pyrolyl)ethyl
2344	benzyl	2-(1-imidazolyl)ethyl

Table 4

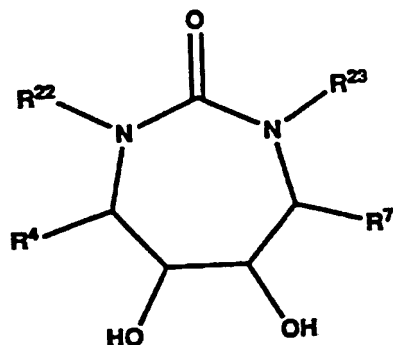
2345	1-pyrolylmethyl	2-(1-imidazolyl) ethyl
2346	1-pyrazolylmethyl	2-(1-imidazolyl) ethyl
2347	1-imidazolylmethyl	2-(1-imidazolyl) ethyl
2348	1-indolylmethyl	2-(1-imidazolyl) ethyl
2349	1-triazolylmethyl	2-(1-imidazolyl) ethyl
2350	1-tetrazolylmethyl	2-(1-imidazolyl) ethyl
2351	2-pyridylmethyl	2-(1-imidazolyl) ethyl
2352	3-pyridylmethyl	2-(1-imidazolyl) ethyl
2353	4-pyridylmethyl	2-(1-imidazolyl) ethyl
2354	cyclohexylmethyl	2-(1-imidazolyl) ethyl
2355	2-naphthylmethyl	2-(1-imidazolyl) ethyl
2356	3-naphthylmethyl	2-(1-imidazolyl) ethyl
2357	2-thiophenylmethyl	2-(1-imidazolyl) ethyl
2358	4-(1-methyl)piperidinyl- methyl	2-(1-imidazolyl) ethyl
2359	"(3,4- methylenedioxyphenyl)methyl"	2-(1-imidazolyl) ethyl
2360	2-thienylmethyl	2-(1-imidazolyl) ethyl
2361	4-biphenylmethyl	2-(1-imidazolyl) ethyl
2362	pyrimidinylmethyl	2-(1-imidazolyl) ethyl
2363	2-benzothiazolylmethyl	2-(1-imidazolyl) ethyl
2364	2-benzothiophenylmethyl	2-(1-imidazolyl) ethyl
2365	2-thiomethylethyl	2-(1-imidazolyl) ethyl
2366	2-thiomethylmethyl	2-(1-imidazolyl) ethyl
2367	2-methylpropyl	2-(1-imidazolyl) ethyl
2368	2-methylbutyl	2-(1-imidazolyl) ethyl
2369	3-methylbutyl	2-(1-imidazolyl) ethyl
2370	cyclopropylmethyl	2-(1-imidazolyl) ethyl
2371	cyclobutylmethyl	2-(1-imidazolyl) ethyl
2372	cyclopentylmethyl	2-(1-imidazolyl) ethyl
2373	p-hydroxyphenylmethyl	2-(1-imidazolyl) ethyl
2374	p-nitrophenylmethyl	2-(1-imidazolyl) ethyl
2375	p-aminophenylmethyl	2-(1-imidazolyl) ethyl
2376	"4-(N,N- dimethylamino)phenylmethyl"	2-(1-imidazolyl) ethyl
2377	benzyl	2-pyridylmethyl
2378	1-pyrolylmethyl	2-pyridylmethyl
2379	1-pyrazolylmethyl	2-pyridylmethyl
2380	1-imidazolylmethyl	2-pyridylmethyl
2381	1-indolylmethyl	2-pyridylmethyl
2382	1-triazolylmethyl	2-pyridylmethyl
2383	1-tetrazolylmethyl	2-pyridylmethyl
2384	2-pyridylmethyl	2-pyridylmethyl
2385	3-pyridylmethyl	2-pyridylmethyl
2386	4-pyridylmethyl	2-pyridylmethyl
2387	cyclohexylmethyl	2-pyridylmethyl
2388	2-naphthylmethyl	2-pyridylmethyl
2389	3-naphthylmethyl	2-pyridylmethyl
2390	2-thiophenylmethyl	2-pyridylmethyl
2391	4-(1-methyl)piperidinyl- methyl	2-pyridylmethyl
2392	"(3,4- methylenedioxyphenyl)methyl"	2-pyridylmethyl
2393	2-thienylmethyl	2-pyridylmethyl
2394	4-biphenylmethyl	2-pyridylmethyl
2395	pyrimidinylmethyl	2-pyridylmethyl
2396	2-benzothiazolylmethyl	2-pyridylmethyl

Table 4

2397	2-benzothiophenylmethyl	2-pyridylmethyl
2398	2-thiomethylethyl	2-pyridylmethyl
2399	2-thiomethylmethyl	2-pyridylmethyl
2400	2-methylpropyl	2-pyridylmethyl
2401	2-methylbutyl	2-pyridylmethyl
2402	3-methylbutyl	2-pyridylmethyl
2403	cyclopropylmethyl	2-pyridylmethyl
2404	cyclobutylmethyl	2-pyridylmethyl
2405	cyclopentylmethyl	2-pyridylmethyl
2406	p-hydroxyphenylmethyl	2-pyridylmethyl
2407	p-nitrophenylmethyl	2-pyridylmethyl
2408	p-aminophenylmethyl	2-pyridylmethyl
2409	"4-(N,N-dimethylamino)phenylmethyl"	2-pyridylmethyl
2410	benzyl	2-thiomethylethyl
2411	1-pyrolylmethyl	2-thiomethylethyl
2412	1-pyrazolylmethyl	2-thiomethylethyl
2413	1-imidazolylmethyl	2-thiomethylethyl
2414	1-indolylmethyl	2-thiomethylethyl
2415	1-triazolylmethyl	2-thiomethylethyl
2416	1-tetrazolylmethyl	2-thiomethylethyl
2417	2-pyridylmethyl	2-thiomethylethyl
2418	3-pyridylmethyl	2-thiomethylethyl
2419	4-pyridylmethyl	2-thiomethylethyl
2420	cyclohexylmethyl	2-thiomethylethyl
2421	2-naphthylmethyl	2-thiomethylethyl
2422	3-naphthylmethyl	2-thiomethylethyl
2423	2-thiophenylmethyl	2-thiomethylethyl
2424	4-(1-methyl)piperidinyl-methyl	2-thiomethylethyl
2425	"(3,4-methylenedioxyphenyl)methyl"	2-thiomethylethyl
2426	2-thienylmethyl	2-thiomethylethyl
2427	4-biphenylmethyl	2-thiomethylethyl
2428	pyrimidinylmethyl	2-thiomethylethyl
2429	2-benzothiazolylmethyl	2-thiomethylethyl
2430	2-benzothiophenylmethyl	2-thiomethylethyl
2431	2-thiomethylethyl	2-thiomethylethyl
2432	2-thiomethylmethyl	2-thiomethylethyl
2433	2-methylpropyl	2-thiomethylethyl
2434	2-methylbutyl	2-thiomethylethyl
2435	3-methylbutyl	2-thiomethylethyl
2436	cyclopropylmethyl	2-thiomethylethyl
2437	cyclobutylmethyl	2-thiomethylethyl
2438	cyclopentylmethyl	2-thiomethylethyl
2439	p-hydroxyphenylmethyl	2-thiomethylethyl
2440	p-nitrophenylmethyl	2-thiomethylethyl
2441	p-aminophenylmethyl	2-thiomethylethyl
2442	"4-(N,N-dimethylamino)phenylmethyl"	2-thiomethylethyl

Table 5

TABLE 5



$$R^4; R^7; R^{22} = R^{23}$$

Example R4
Number

R7

R22 = R23

Example R4 Number		R7	R22 = R23
2443	benzyl	"4-(N,N-dimethylamino)phenyl-methyl"	allyl
2444	1-pyrrolylmethyl	benzyl	allyl
2445	1-pyrazolylmethyl	1-pyrrolylmethyl	allyl
2446	1-imidazolylmethyl	1-pyrazolylmethyl	allyl
2447	1-indolylmethyl	1-imidazolylmethyl	allyl
2448	1-triazolylmethyl	1-indolylmethyl	allyl
2449	1-tetrazolylmethyl	1-triazolylmethyl	allyl
2450	2-pyridylmethyl	1-tetrazolylmethyl	allyl
2451	3-pyridylmethyl	2-pyridylmethyl	allyl
2452	4-pyridylmethyl	3-pyridylmethyl	allyl
2453	cyclohexylmethyl	4-pyridylmethyl	allyl
2454	2-naphthylmethyl	cyclohexylmethyl	allyl
2455	3-naphthylmethyl	2-naphthylmethyl	allyl
2456	2-thiophenylmethyl	3-naphthylmethyl	allyl
2457	4-(1-methyl)piperidinylmethyl	2-thiophenylmethyl	allyl
2458	"(3,4-methylenedioxy-phenyl)methyl"	4-(1-methyl)piperidinyl-methyl	allyl
2459	2-thienylmethyl	"(3,4-methylenedioxy-phenyl)methyl"	allyl
2460	4-biphenylmethyl	2-thienylmethyl	allyl
2461	pyrimidinylmethyl	4-biphenylmethyl	allyl
2462	2-benzothiazolyl-methyl	pyrimidinylmethyl	allyl
2463	2-benzothiophenyl-methyl	2-benzothiazolyl-methyl	allyl

Table 5

2464	2-thiomethylethyl	2-benzothiophenyl-methyl	allyl
2465	2-thiomethylmethyl	2-thiomethylethyl	allyl
2466	2-methylpropyl	2-thiomethylmethyl	allyl
2467	2-methylbutyl	2-methylpropyl	allyl
2468	3-methylbutyl	2-methylbutyl	allyl
2469	cyclopropylmethyl	3-methylbutyl	allyl
2470	cyclobutylmethyl	cyclopropylmethyl	allyl
2471	cyclopentylmethyl	cyclobutylmethyl	allyl
2472	p-hydroxyphenyl-methyl	cyclopentylmethyl	allyl
2473	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	allyl
2474	p-aminophenyl-methyl	p-nitrophenylmethyl	allyl
2475	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	allyl
2476	benzyl	"4-(N,N-dimethylamino)-phenyl-methyl"	propyl
2477	1-pyrollylmethyl	benzyl	propyl
2478	1-pyrazolylmethyl	1-pyrollylmethyl	propyl
2479	1-imidazolylmethyl	1-pyrazolylmethyl	propyl
2480	1-indolylmethyl	1-imidazolylmethyl	propyl
2481	1-triazolylmethyl	1-indolylmethyl	propyl
2482	1-tetrazolylmethyl	1-triazolylmethyl	propyl
2483	2-pyridylmethyl	1-tetrazolylmethyl	propyl
2484	3-pyridylmethyl	2-pyridylmethyl	propyl
2485	4-pyridylmethyl	3-pyridylmethyl	propyl
2486	cyclohexylmethyl	4-pyridylmethyl	propyl
2487	2-naphthylmethyl	cyclohexylmethyl	propyl
2488	3-naphthylmethyl	2-naphthylmethyl	propyl
2489	2-thiophenylmethyl	3-naphthylmethyl	propyl
2490	4-(1-methyl)-piperidinyl-methyl	2-thiophenylmethyl	propyl
2491	"(3,4-methylene-dioxyphenyl)methyl"	4-(1-methyl)-piperidinyl-methyl	propyl
2492	2-thienylmethyl	"(3,4-methylene-dioxyphenyl)methyl"	propyl
2493	4-biphenylmethyl	2-thienylmethyl	propyl
2494	pyrimidinylmethyl	4-biphenylmethyl	propyl
2495	2-benzothiazolylmethyl	pyrimidinylmethyl	propyl
2496	2-benzothiophenylmethyl	2-benzothiazolylmethyl	propyl
2497	2-thiomethylethyl	2-benzothiophenylmethyl	propyl
2498	2-thiomethylmethyl	2-thiomethylethyl	propyl
2499	2-methylpropyl	2-thiomethylmethyl	propyl
2500	2-methylbutyl	2-methylpropyl	propyl
2501	3-methylbutyl	2-methylbutyl	propyl
2502	cyclopropylmethyl	3-methylbutyl	propyl
2503	cyclobutylmethyl	cyclopropylmethyl	propyl
2504	cyclopentylmethyl	cyclobutylmethyl	propyl
2505	p-hydroxyphenyl-methyl	cyclopentylmethyl	propyl
2506	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	propyl
2507	p-aminophenyl-methyl	p-nitrophenylmethyl	propyl
2508	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	propyl
2509	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	n-butyl
2510	1-pyrollylmethyl	benzyl	n-butyl
2511	1-pyrazolylmethyl	1-pyrollylmethyl	n-butyl

Table 5

2512	1-imidazolylmethyl	1-pyrazolylmethyl	n-butyl
2513	1-indolylmethyl	1-imidazolylmethyl	n-butyl
2514	1-triazolylmethyl	1-indolylmethyl	n-butyl
2515	1-tetrazolylmethyl	1-triazolylmethyl	n-butyl
2516	2-pyridylmethyl	1-tetrazolylmethyl	n-butyl
2517	3-pyridylmethyl	2-pyridylmethyl	n-butyl
2518	4-pyridylmethyl	3-pyridylmethyl	n-butyl
2519	cyclohexylmethyl	4-pyridylmethyl	n-butyl
2520	2-naphthylmethyl	cyclohexylmethyl	n-butyl
2521	3-naphthylmethyl	2-naphthylmethyl	n-butyl
2522	2-thiophenylmethyl	3-naphthylmethyl	n-butyl
2523	4-(1-methyl)- piperidinyl-methyl	2-thiophenylmethyl	n-butyl
2524	"(3,4- methylenedioxyphenyl)m ethyl"	4-(1-methyl)- piperidinyl-methyl	n-butyl
2525	2-thienylmethyl	"(3,4-methylene- dioxyphenyl)methyl"	n-butyl
2526	4-biphenylmethyl	2-thienylmethyl	n-butyl
2527	pyrimidinylmethyl	4-biphenylmethyl	n-butyl
2528	2-benzothiazolylmethyl	pyrimidinylmethyl	n-butyl
2529	2- benzothiophenylmethyl	2-benzothiazolylmethyl	n-butyl
2530	2-thiomethylethyl	2- benzothiophenylmethyl	n-butyl
2531	2-thiomethylmethyl	2-thiomethylethyl	n-butyl
2532	2-methylpropyl	2-thiomethylmethyl	n-butyl
2533	2-methylbutyl	2-methylpropyl	n-butyl
2534	3-methylbutyl	2-methylbutyl	n-butyl
2535	cyclopropylmethyl	3-methylbutyl	n-butyl
2536	cyclobutylmethyl	cyclopropylmethyl	n-butyl
2537	cyclopentylmethyl	cyclobutylmethyl	n-butyl
2538	p-hydroxyphenyl-methyl	cyclopentylmethyl	n-butyl
2539	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	n-butyl
2540	p-aminophenyl-methyl	p-nitrophenylmethyl	n-butyl
2541	"4-(N,N-dimethyl- amino)phenyl-methyl"	p-aminophenyl-methyl	n-butyl
2542	benzyl	"4-(N,N-dimethyl- amino)-phenylmethyl"	isobutyl
2543	1-pyrollylmethyl	benzyl	isobutyl
2544	1-pyrazolylmethyl	1-pyrollylmethyl	isobutyl
2545	1-imidazolylmethyl	1-pyrazolylmethyl	isobutyl
2546	1-indolylmethyl	1-imidazolylmethyl	isobutyl
2547	1-triazolylmethyl	1-indolylmethyl	isobutyl
2548	1-tetrazolylmethyl	1-triazolylmethyl	isobutyl
2549	2-pyridylmethyl	1-tetrazolylmethyl	isobutyl
2550	3-pyridylmethyl	2-pyridylmethyl	isobutyl
2551	4-pyridylmethyl	3-pyridylmethyl	isobutyl
2552	cyclohexylmethyl	4-pyridylmethyl	isobutyl
2553	2-naphthylmethyl	cyclohexylmethyl	isobutyl
2554	3-naphthylmethyl	2-naphthylmethyl	isobutyl
2555	2-thiophenylmethyl	3-naphthylmethyl	isobutyl
2556	4-(1-methyl)- piperidinyl-methyl	2-thiophenylmethyl	isobutyl
2557	"(3,4- methylenedioxyphenyl)m ethyl"	4-(1-methyl)- piperidinyl-methyl	isobutyl

Table 5

2558	2-thienylmethyl	"(3,4-methylene-dioxyphenyl)methyl"	isobutyl
2559	4-biphenylmethyl	2-thienylmethyl	isobutyl
2560	pyrimidinylmethyl	4-biphenylmethyl	isobutyl
2561	2-benzothiazolylmethyl	pyrimidinylmethyl	isobutyl
2562	2-benzothiophenylmethyl	2-benzothiazolylmethyl	isobutyl
2563	2-thiomethylethyl	2-benzothiophenylmethyl	isobutyl
2564	2-thiomethylmethyl	2-thiomethylethyl	isobutyl
2565	2-methylpropyl	2-thiomethylmethyl	isobutyl
2566	2-methylbutyl	2-methylpropyl	isobutyl
2567	3-methylbutyl	2-methylbutyl	isobutyl
2568	cyclopropylmethyl	3-methylbutyl	isobutyl
2569	cyclobutylmethyl	cyclopropylmethyl	isobutyl
2570	cyclopentylmethyl	cyclobutylmethyl	isobutyl
2571	p-hydroxyphenyl-methyl	cyclopentylmethyl	isobutyl
2572	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	isobutyl
2573	p-aminophenyl-methyl	p-nitrophenylmethyl	isobutyl
2574	"4-(N,N-dimethyl-amino)-phenylmethyl"	p-aminophenyl-methyl	isobutyl
2575	benzyl	"4-(N,N-dimethyl-amino)-phenylmethyl"	2-butyl
2576	1-pyrollylmethyl	benzyl	2-butyl
2577	1-pyrazolylmethyl	1-pyrollylmethyl	2-butyl
2578	1-imidazolylmethyl	1-pyrazolylmethyl	2-butyl
2579	1-indolylmethyl	1-imidazolylmethyl	2-butyl
2580	1-triazolylmethyl	1-indolylmethyl	2-butyl
2581	1-tetrazolylmethyl	1-triazolylmethyl	2-butyl
2582	2-pyridylmethyl	1-tetrazolylmethyl	2-butyl
2583	3-pyridylmethyl	2-pyridylmethyl	2-butyl
2584	4-pyridylmethyl	3-pyridylmethyl	2-butyl
2585	cyclohexylmethyl	4-pyridylmethyl	2-butyl
2586	2-naphthylmethyl	cyclohexylmethyl	2-butyl
2587	3-naphthylmethyl	2-naphthylmethyl	2-butyl
2588	2-thiophenylmethyl	3-naphthylmethyl	2-butyl
2589	4-(1-methyl)-piperidinyl-methyl	2-thiophenylmethyl	2-butyl
2590	"(3,4-methylene-dioxyphenyl)methyl"	4-(1-methyl)-piperidinyl-methyl	2-butyl
2591	2-thienylmethyl	"(3,4-methylene-dioxyphenyl)methyl"	2-butyl
2592	4-biphenylmethyl	2-thienylmethyl	2-butyl
2593	pyrimidinylmethyl	4-biphenylmethyl	2-butyl
2594	2-benzothiazolylmethyl	pyrimidinylmethyl	2-butyl
2595	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-butyl
2596	2-thiomethylethyl	2-benzothiophenylmethyl	2-butyl
2597	2-thiomethylmethyl	2-thiomethylethyl	2-butyl
2598	2-methylpropyl	2-thiomethylmethyl	2-butyl
2599	2-methylbutyl	2-methylpropyl	2-butyl
2600	3-methylbutyl	2-methylbutyl	2-butyl
2601	cyclopropylmethyl	3-methylbutyl	2-butyl
2602	cyclobutylmethyl	cyclopropylmethyl	2-butyl
2603	cyclopentylmethyl	cyclobutylmethyl	2-butyl
2604	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-butyl
2605	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-butyl

Table 5

2606	p-aminophenyl-methyl	p-nitrophenylmethyl	2-butyl	
2607	"4-(N,N-dimethyl- amino)-phenylmethyl"	p-aminophenyl-methyl	2-butyl	
2608	benzyl	"4-(N,N-dimethyl- amino) phenylmethyl"	"3,3- dimethallyl"	
2609	1-pyrolylmethyl	benzyl	"3,3- dimethallyl"	
2610	1-pyrazolylmethyl	1-pyrolylmethyl	"3,3- dimethallyl"	
2611	1-imidazolylmethyl	1-pyrazolylmethyl	"3,3- dimethallyl"	
2612	1-indolylmethyl	1-imidazolylmethyl	"3,3- dimethallyl"	
2613	1-triazolylmethyl	1-indolylmethyl	"3,3- dimethallyl"	
2614	1-tetrazolylmethyl	1-triazolylmethyl	"3,3- dimethallyl"	
2615	2-pyridylmethyl	1-tetrazolylmethyl	"3,3- dimethallyl"	
2616	3-pyridylmethyl	2-pyridylmethyl	"3,3- dimethallyl"	
2617	4-pyridylmethyl	3-pyridylmethyl	"3,3- dimethallyl"	
2618	cyclohexylmethyl	4-pyridylmethyl	"3,3- dimethallyl"	
2619	2-naphthylmethyl	cyclohexylmethyl	"3,3- dimethallyl"	
2620	3-naphthylmethyl	2-naphthylmethyl	"3,3- dimethallyl"	
2621	2-thiophenylmethyl	3-naphthylmethyl	"3,3- dimethallyl"	
2622	4-(1- methyl)piperidinyl- methyl	2-thiophenylmethyl	"3,3- dimethallyl"	
2623	"(3,4- methylenedioxyphenyl)m ethyl"	4-(1- methyl)piperidinyl- methyl	"3,3- dimethallyl"	
2624	2-thienylmethyl	"(3,4-methylene- dioxyphenyl)methyl"	"3,3- dimethallyl"	
2625	4-biphenylmethyl	2-thienylmethyl	"3,3- dimethallyl"	
2626	pyrimidinylmethyl	4-biphenylmethyl	"3,3- dimethallyl"	
2627	2-benzothiazolylmethyl	pyrimidinylmethyl	"3,3- dimethallyl"	
2628	2- benzothiophenylmethyl	2-benzothiazolylmethyl	"3,3- dimethallyl"	
2629	2-thiomethylethyl	2- benzothiophenylmethyl	"3,3- dimethallyl"	
2630	2-thiomethylmethyl	2-thiomethylethyl	"3,3- dimethallyl"	
2631	2-methylpropyl	2-thiomethylmethyl	"3,3- dimethallyl"	
2632	2-methylbutyl	2-methylpropyl	"3,3- dimethallyl"	
2633	3-methylbutyl	2-methylbutyl	"3,3- dimethallyl"	

Table 5

2634	cyclopropylmethyl	3-methylbutyl	"3,3-dimethyl"
2635	cyclobutylmethyl	cyclopropylmethyl	"3,3-dimethyl"
2636	cyclopentylmethyl	cyclobutylmethyl	"3,3-dimethyl"
2637	p-hydroxyphenyl-methyl	cyclopentylmethyl	"3,3-dimethyl"
2638	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	"3,3-dimethyl"
2639	p-aminophenyl-methyl	p-nitrophenylmethyl	"3,3-dimethyl"
2640	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	"3,3-dimethyl"
2641	benzyl	"4-(N,N-dimethylamino)-phenyl-methyl"	3-methyl
2642	1-pyrollylmethyl	benzyl	3-methyl
2643	1-pyrazolylmethyl	1-pyrollylmethyl	3-methyl
2644	1-imidazolylmethyl	1-pyrazolylmethyl	3-methyl
2645	1-indolylmethyl	1-imidazolylmethyl	3-methyl
2646	1-triazolylmethyl	1-indolylmethyl	3-methyl
2647	1-tetrazolylmethyl	1-triazolylmethyl	3-methyl
2648	2-pyridylmethyl	1-tetrazolylmethyl	3-methyl
2649	3-pyridylmethyl	2-pyridylmethyl	3-methyl
2650	4-pyridylmethyl	3-pyridylmethyl	3-methyl
2651	cyclohexylmethyl	4-pyridylmethyl	3-methyl
2652	2-naphthylmethyl	cyclohexylmethyl	3-methyl
2653	3-naphthylmethyl	2-naphthylmethyl	3-methyl
2654	2-thiophenylmethyl	3-naphthylmethyl	3-methyl
2655	4-(1-methyl)-piperidinyl-methyl	2-thiophenylmethyl	3-methyl
2656	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	3-methyl
2657	2-thienylmethyl	"(3,4-methylene-dioxyphenyl)methyl"	3-methyl
2658	4-biphenylmethyl	2-thienylmethyl	3-methyl
2659	pyrimidinylmethyl	4-biphenylmethyl	3-methyl
2660	2-benzothiazolylmethyl	pyrimidinylmethyl	3-methyl
2661	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-methyl
2662	2-thiomethylethyl	2-benzothiophenylmethyl	3-methyl
2663	2-thiomethylmethyl	2-thiomethylethyl	3-methyl
2664	2-methylpropyl	2-thiomethylmethyl	3-methyl
2665	2-methylbutyl	2-methylpropyl	3-methyl
2666	3-methylbutyl	2-methylbutyl	3-methyl
2667	cyclopropylmethyl	3-methylbutyl	3-methyl
2668	cyclobutylmethyl	cyclopropylmethyl	3-methyl
2669	cyclopentylmethyl	cyclobutylmethyl	3-methyl
2670	p-hydroxyphenyl-methyl	cyclopentylmethyl	3-methyl
2671	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	3-methyl
2672	p-aminophenyl-methyl	p-nitrophenylmethyl	3-methyl
2673	"4-(N,N-dimethyl-amino)-phenyl-methyl"	p-aminophenyl-methyl	3-methyl

Table 5

2674	benzyl	"4-(N,N-dimethyl-amino)phenylmethyl"	2-methallyl
2675	1-pyrollylmethyl	benzyl	2-methallyl
2676	1-pyrazolylmethyl	1-pyrollylmethyl	2-methallyl
2677	1-imidazolylmethyl	1-pyrazolylmethyl	2-methallyl
2678	1-indolylmethyl	1-imidazolylmethyl	2-methallyl
2679	1-triazolylmethyl	1-indolylmethyl	2-methallyl
2680	1-tetrazolylmethyl	1-triazolylmethyl	2-methallyl
2681	2-pyridylmethyl	1-tetrazolylmethyl	2-methallyl
2682	3-pyridylmethyl	2-pyridylmethyl	2-methallyl
2683	4-pyridylmethyl	3-pyridylmethyl	2-methallyl
2684	cyclohexylmethyl	4-pyridylmethyl	2-methallyl
2685	2-naphthylmethyl	cyclohexylmethyl	2-methallyl
2686	3-naphthylmethyl	2-naphthylmethyl	2-methallyl
2687	2-thiophenylmethyl	3-naphthylmethyl	2-methallyl
2688	4-(1-methyl)piperidinylmethyl	2-thiophenylmethyl	2-methallyl
2689	"(3,4-methylene-dioxyphenyl)methyl"	4-(1-methyl)-piperidinyl-methyl	2-methallyl
2690	2-thienylmethyl	"(3,4-methylene-dioxyphenyl)methyl"	2-methallyl
2691	4-biphenylmethyl	2-thienylmethyl	2-methallyl
2692	pyrimidinylmethyl	4-biphenylmethyl	2-methallyl
2693	2-benzothiazolylmethyl	pyrimidinylmethyl	2-methallyl
2694	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-methallyl
2695	2-thiomethylethyl	2-benzothiophenylmethyl	2-methallyl
2696	2-thiomethylmethyl	2-thiomethylethyl	2-methallyl
2697	2-methylpropyl	2-thiomethylmethyl	2-methallyl
2698	2-methylbutyl	2-methylpropyl	2-methallyl
2699	3-methylbutyl	2-methylbutyl	2-methallyl
2700	cyclopropylmethyl	3-methylbutyl	2-methallyl
2701	cyclobutylmethyl	cyclopropylmethyl	2-methallyl
2702	cyclopentylmethyl	cyclobutylmethyl	2-methallyl
2703	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-methallyl
2704	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-methallyl
2705	p-aminophenyl-methyl	p-nitrophenylmethyl	2-methallyl
2706	"4-(N,N-dimethyl-amino)-phenylmethyl"	p-aminophenyl-methyl	2-methallyl
2707	benzyl	"4-(N,N-dimethyl-amino)-phenylmethyl"	2-propyl
2708	1-pyrollylmethyl	benzyl	2-propyl
2709	1-pyrazolylmethyl	1-pyrollylmethyl	2-propyl
2710	1-imidazolylmethyl	1-pyrazolylmethyl	2-propyl
2711	1-indolylmethyl	1-imidazolylmethyl	2-propyl
2712	1-triazolylmethyl	1-indolylmethyl	2-propyl
2713	1-tetrazolylmethyl	1-triazolylmethyl	2-propyl
2714	2-pyridylmethyl	1-tetrazolylmethyl	2-propyl
2715	3-pyridylmethyl	2-pyridylmethyl	2-propyl
2716	4-pyridylmethyl	3-pyridylmethyl	2-propyl
2717	cyclohexylmethyl	4-pyridylmethyl	2-propyl
2718	2-naphthylmethyl	cyclohexylmethyl	2-propyl
2719	3-naphthylmethyl	2-naphthylmethyl	2-propyl
2720	2-thiophenylmethyl	3-naphthylmethyl	2-propyl

Table 5

2721	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-propyl
2722	"(3,4-methylene-dioxyphenyl)methyl"	4-(1-methyl)-piperidinyl-methyl	2-propyl
2723	2-thienylmethyl	"(3,4-methylene-dioxyphenyl)methyl"	2-propyl
2724	4-biphenylmethyl	2-thienylmethyl	2-propyl
2725	pyrimidinylmethyl	4-biphenylmethyl	2-propyl
2726	2-benzothiazolylmethyl	pyrimidinylmethyl	2-propyl
2727	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-propyl
2728	2-thiomethylethyl	2-benzothiophenylmethyl	2-propyl
2729	2-thiomethylmethyl	2-thiomethylethyl	2-propyl
2730	2-methylpropyl	2-thiomethylmethyl	2-propyl
2731	2-methylbutyl	2-methylpropyl	2-propyl
2732	3-methylbutyl	2-methylbutyl	2-propyl
2733	cyclopropylmethyl	3-methylbutyl	2-propyl
2734	cyclobutylmethyl	cyclopropylmethyl	2-propyl
2735	cyclopentylmethyl	cyclobutylmethyl	2-propyl
2736	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-propyl
2737	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-propyl
2738	p-aminophenyl-methyl	p-nitrophenylmethyl	2-propyl
2739	"4-(N,N-dimethyl-amino)-phenylmethyl"	p-aminophenyl-methyl	2-propyl
2740	benzyl	"4-(N,N-dimethyl-amino)phenylmethyl"	cyclopropyl
2741	1-pyrollylmethyl	benzyl	cyclopropyl
2742	1-pyrazolylmethyl	1-pyrollylmethyl	cyclopropyl
2743	1-imidazolylmethyl	1-pyrazolylmethyl	cyclopropyl
2744	1-indolylmethyl	1-imidazolylmethyl	cyclopropyl
2745	1-triazolylmethyl	1-indolylmethyl	cyclopropyl
2746	1-tetrazolylmethyl	1-triazolylmethyl	cyclopropyl
2747	2-pyridylmethyl	1-tetrazolylmethyl	cyclopropyl
2748	3-pyridylmethyl	2-pyridylmethyl	cyclopropyl
2749	4-pyridylmethyl	3-pyridylmethyl	cyclopropyl
2750	cyclohexylmethyl	4-pyridylmethyl	cyclopropyl
2751	2-naphthylmethyl	cyclohexylmethyl	cyclopropyl
2752	3-naphthylmethyl	2-naphthylmethyl	cyclopropyl
2753	2-thiophenylmethyl	3-naphthylmethyl	cyclopropyl
2754	4-(1-methyl)-piperidinyl-methyl	2-thiophenylmethyl	cyclopropyl
2755	"(3,4-methylene-dioxyphenyl)methyl"	4-(1-methyl)-piperidinyl-methyl	cyclopropyl
2756	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclopropyl
2757	4-biphenylmethyl	2-thienylmethyl	cyclopropyl
2758	pyrimidinylmethyl	4-biphenylmethyl	cyclopropyl
2759	2-benzothiazolylmethyl	pyrimidinylmethyl	cyclopropyl
2760	2-benzothiophenylmethyl	2-benzothiazolylmethyl	cyclopropyl
2761	2-thiomethylethyl	2-benzothiophenylmethyl	cyclopropyl
2762	2-thiomethylmethyl	2-thiomethylethyl	cyclopropyl
2763	2-methylpropyl	2-thiomethylmethyl	cyclopropyl
2764	2-methylbutyl	2-methylpropyl	cyclopropyl

Table 5

2765	3-methylbutyl	2-methylbutyl	cyclopropyl
2766	cyclopropylmethyl	3-methylbutyl	cyclopropyl
2767	cyclobutylmethyl	cyclopropylmethyl	cyclopropyl
2768	cyclopentylmethyl	cyclobutylmethyl	cyclopropyl
2769	p-hydroxyphenyl-methyl	cyclopentylmethyl	cyclopropyl
2770	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	cyclopropyl
2771	p-aminophenyl-methyl	p-nitrophenylmethyl	cyclopropyl
2772	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	cyclopropyl
2773	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	cyclopropylmethyl
2774	1-pyrrolylmethyl	benzyl	hyl
2775	1-pyrazolylmethyl	1-pyrrolylmethyl	cyclopropylmethyl
2776	1-imidazolylmethyl	1-pyrazolylmethyl	hyl
2777	1-indolylmethyl	1-imidazolylmethyl	cyclopropylmethyl
2778	1-triazolylmethyl	1-indolylmethyl	hyl
2779	1-tetrazolylmethyl	1-triazolylmethyl	cyclopropylmethyl
2780	2-pyridylmethyl	1-tetrazolylmethyl	hyl
2781	3-pyridylmethyl	2-pyridylmethyl	cyclopropylmethyl
2782	4-pyridylmethyl	3-pyridylmethyl	hyl
2783	cyclohexylmethyl	4-pyridylmethyl	cyclopropylmethyl
2784	2-naphthylmethyl	cyclohexylmethyl	hyl
2785	3-naphthylmethyl	2-naphthylmethyl	cyclopropylmethyl
2786	2-thiophenylmethyl	3-naphthylmethyl	hyl
2787	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	cyclopropylmethyl
2788	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	hyl
2789	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclopropylmethyl
2790	4-biphenylmethyl	2-thienylmethyl	hyl
2791	pyrimidinylmethyl	4-biphenylmethyl	cyclopropylmethyl
2792	2-benzothiazolylmethyl	pyrimidinylmethyl	hyl
2793	2-benzothiophenylmethyl	2-benzothiazolylmethyl	cyclopropylmethyl
2794	2-thiomethylethyl	2-benzothiophenylmethyl	hyl

Table 5

2795	2-thiomethylmethyl	2-thiomethylethyl	cyclopropylmet hyl
2796	2-methylpropyl	2-thiomethylmethyl	cyclopropylmet hyl
2797	2-methylbutyl	2-methylpropyl	cyclopropylmet hyl
2798	3-methylbutyl	2-methylbutyl	cyclopropylmet hyl
2799	cyclopropylmethyl	3-methylbutyl	cyclopropylmet hyl
2800	cyclobutylmethyl	cyclopropylmethyl	cyclopropylmet hyl
2801	cyclopentylmethyl	cyclobutylmethyl	cyclopropylmet hyl
2802	p-hydroxyphenyl-methyl	cyclopentylmethyl	cyclopropylmet hyl
2803	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	cyclopropylmet hyl
2804	p-aminophenyl-methyl	p-nitrophenylmethyl	cyclopropylmet hyl
2805	"4-(N,N- dimethylamino)phenylme thyl"	p-aminophenyl-methyl	cyclopropylmet hyl
2806	benzyl	"4-(N,N- dimethylamino)phenylme thyl"	n-pentyl
2807	1-pyrolylmethyl	benzyl	n-pentyl
2808	1-pyrazolylmethyl	1-pyrolylmethyl	n-pentyl
2809	1-imidazolylmethyl	1-pyrazolylmethyl	n-pentyl
2810	1-indolylmethyl	1-imidazolylmethyl	n-pentyl
2811	1-triazolylmethyl	1-indolylmethyl	n-pentyl
2812	1-tetrazolylmethyl	1-triazolylmethyl	n-pentyl
2813	2-pyridylmethyl	1-tetrazolylmethyl	n-pentyl
2814	3-pyridylmethyl	2-pyridylmethyl	n-pentyl
2815	4-pyridylmethyl	3-pyridylmethyl	n-pentyl
2816	cyclohexylmethyl	4-pyridylmethyl	n-pentyl
2817	2-naphthylmethyl	cyclohexylmethyl	n-pentyl
2818	3-naphthylmethyl	2-naphthylmethyl	n-pentyl
2819	2-thiophenylmethyl	3-naphthylmethyl	n-pentyl
2820	4-(1- methyl)piperidinyl- methyl	2-thiophenylmethyl	n-pentyl
2821	"(3,4- methylenedioxyphenyl)m ethyl"	4-(1- methyl)piperidinyl- methyl	n-pentyl
2822	2-thienylmethyl	"(3,4- methylenedioxyphenyl)m ethyl"	n-pentyl
2823	4-biphenylmethyl	2-thienylmethyl	n-pentyl
2824	pyrimidinylmethyl	4-biphenylmethyl	n-pentyl
2825	2-benzothiazolylmethyl	pyrimidinylmethyl	n-pentyl
2826	2- benzothiophenylmethyl	2-benzothiazolylmethyl	n-pentyl
2827	2-thiomethylethyl	2- benzothiophenylmethyl	n-pentyl
2828	2-thiomethylmethyl	2-thiomethylethyl	n-pentyl
2829	2-methylpropyl	2-thiomethylmethyl	n-pentyl
2830	2-methylbutyl	2-methylpropyl	n-pentyl

Table 5

2831	3-methylbutyl	2-methylbutyl	n-pentyl
2832	cyclopropylmethyl	3-methylbutyl	n-pentyl
2833	cyclobutylmethyl	cyclopropylmethyl	n-pentyl
2834	cyclopentylmethyl	cyclobutylmethyl	n-pentyl
2835	p-hydroxyphenyl-methyl	cyclopentylmethyl	n-pentyl
2836	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	n-pentyl
2837	p-aminophenyl-methyl	p-nitrophenylmethyl	n-pentyl
2838	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	n-pentyl
2839	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-pentyl
2840	1-pyrrolylmethyl	benzyl	2-pentyl
2841	1-pyrazolylmethyl	1-pyrrolylmethyl	2-pentyl
2842	1-imidazolylmethyl	1-pyrazolylmethyl	2-pentyl
2843	1-indolylmethyl	1-imidazolylmethyl	2-pentyl
2844	1-triazolylmethyl	1-indolylmethyl	2-pentyl
2845	1-tetrazolylmethyl	1-triazolylmethyl	2-pentyl
2846	2-pyridylmethyl	1-tetrazolylmethyl	2-pentyl
2847	3-pyridylmethyl	2-pyridylmethyl	2-pentyl
2848	4-pyridylmethyl	3-pyridylmethyl	2-pentyl
2849	cyclohexylmethyl	4-pyridylmethyl	2-pentyl
2850	2-naphthylmethyl	cyclohexylmethyl	2-pentyl
2851	3-naphthylmethyl	2-naphthylmethyl	2-pentyl
2852	2-thiophenylmethyl	3-naphthylmethyl	2-pentyl
2853	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-pentyl
2854	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-pentyl
2855	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-pentyl
2856	4-biphenylmethyl	2-thienylmethyl	2-pentyl
2857	pyrimidinylmethyl	4-biphenylmethyl	2-pentyl
2858	2-benzothiazolylmethyl	pyrimidinylmethyl	2-pentyl
2859	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-pentyl
2860	2-thiomethylethyl	2-benzothiophenylmethyl	2-pentyl
2861	2-thiomethylmethyl	2-thiomethylethyl	2-pentyl
2862	2-methylpropyl	2-thiomethylmethyl	2-pentyl
2863	2-methylbutyl	2-methylpropyl	2-pentyl
2864	3-methylbutyl	2-methylbutyl	2-pentyl
2865	cyclopropylmethyl	3-methylbutyl	2-pentyl
2866	cyclobutylmethyl	cyclopropylmethyl	2-pentyl
2867	cyclopentylmethyl	cyclobutylmethyl	2-pentyl
2868	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-pentyl
2869	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-pentyl
2870	p-aminophenyl-methyl	p-nitrophenylmethyl	2-pentyl
2871	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-pentyl
2872	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	3-pentyl

Table 5

2873	1-pyrolylmethyl	benzyl	3-pentyl
2874	1-pyrazolylmethyl	1-pyrolylmethyl	3-pentyl
2875	1-imidazolylmethyl	1-pyrazolylmethyl	3-pentyl
2876	1-indolylmethyl	1-imidazolylmethyl	3-pentyl
2877	1-triazolylmethyl	1-indolylmethyl	3-pentyl
2878	1-tetrazolylmethyl	1-triazolylmethyl	3-pentyl
2879	2-pyridylmethyl	1-tetrazolylmethyl	3-pentyl
2880	3-pyridylmethyl	2-pyridylmethyl	3-pentyl
2881	4-pyridylmethyl	3-pyridylmethyl	3-pentyl
2882	cyclohexylmethyl	4-pyridylmethyl	3-pentyl
2883	2-naphthylmethyl	cyclohexylmethyl	3-pentyl
2884	3-naphthylmethyl	2-naphthylmethyl	3-pentyl
2885	2-thiophenylmethyl	3-naphthylmethyl	3-pentyl
2886	4-(1-methyl)piperidinyl-	2-thiophenylmethyl	3-pentyl
	methyl		
2887	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-	3-pentyl
	ethyl"	methyl	
2888	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-pentyl
2889	4-biphenylmethyl	2-thienylmethyl	3-pentyl
2890	pyrimidinylmethyl	4-biphenylmethyl	3-pentyl
2891	2-benzothiazolylmethyl	pyrimidinylmethyl	3-pentyl
2892	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-pentyl
2893	2-thiomethylethyl	2-benzothiophenylmethyl	3-pentyl
2894	2-thiomethylmethyl	2-thiomethylethyl	3-pentyl
2895	2-methylpropyl	2-thiomethylmethyl	3-pentyl
2896	2-methylbutyl	2-methylpropyl	3-pentyl
2897	3-methylbutyl	2-methylbutyl	3-pentyl
2898	cyclopropylmethyl	3-methylbutyl	3-pentyl
2899	cyclobutylmethyl	cyclopropylmethyl	3-pentyl
2900	cyclopentylmethyl	cyclobutylmethyl	3-pentyl
2901	p-hydroxyphenyl-methyl	cyclopentylmethyl	3-pentyl
2902	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	3-pentyl
2903	p-aminophenyl-methyl	p-nitrophenylmethyl	3-pentyl
2904	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	3-pentyl
	benzyl		
2905		"4-(N,N-dimethylamino)phenylmethyl"	3-methylbutyl
		benzyl	
2906	1-pyrolylmethyl	1-pyrolylmethyl	3-methylbutyl
2907	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methylbutyl
2908	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl
2909	1-indolylmethyl	1-indolylmethyl	3-methylbutyl
2910	1-triazolylmethyl	1-triazolylmethyl	3-methylbutyl
2911	1-tetrazolylmethyl	1-tetrazolylmethyl	3-methylbutyl
2912	2-pyridylmethyl	2-pyridylmethyl	3-methylbutyl
2913	3-pyridylmethyl	3-pyridylmethyl	3-methylbutyl
2914	4-pyridylmethyl	4-pyridylmethyl	3-methylbutyl
2915	cyclohexylmethyl	cyclohexylmethyl	3-methylbutyl
2916	2-naphthylmethyl	2-naphthylmethyl	3-methylbutyl
2917	3-naphthylmethyl	3-naphthylmethyl	3-methylbutyl
2918	2-thiophenylmethyl		

Table 5

2919	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	3-methylbutyl
2920	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	3-methylbutyl
2921	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-methylbutyl
2922	4-biphenylmethyl	2-thienylmethyl	3-methylbutyl
2923	pyrimidinylmethyl	4-biphenylmethyl	3-methylbutyl
2924	2-benzothiazolylmethyl	pyrimidinylmethyl	3-methylbutyl
2925	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-methylbutyl
2926	2-thiomethylethyl	2-benzothiophenylmethyl	3-methylbutyl
2927	2-thiomethylmethyl	2-thiomethylethyl	3-methylbutyl
2928	2-methylpropyl	2-thiomethylmethyl	3-methylbutyl
2929	2-methylbutyl	2-methylpropyl	3-methylbutyl
2930	3-methylbutyl	2-methylbutyl	3-methylbutyl
2931	cyclopropylmethyl	3-methylbutyl	3-methylbutyl
2932	cyclobutylmethyl	cyclopropylmethyl	3-methylbutyl
2933	cyclopentylmethyl	cyclobutylmethyl	3-methylbutyl
2934	p-hydroxyphenyl-methyl	cyclopentylmethyl	3-methylbutyl
2935	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	3-methylbutyl
2936	p-aminophenyl-methyl	p-nitrophenylmethyl	3-methylbutyl
2937	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	3-methylbutyl
2938	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-methylbutyl
2939	1-pyrollylmethyl	benzyl	2-methylbutyl
2940	1-pyrazolylmethyl	1-pyrollylmethyl	2-methylbutyl
2941	1-imidazolylmethyl	1-pyrazolylmethyl	2-methylbutyl
2942	1-indolylmethyl	1-imidazolylmethyl	2-methylbutyl
2943	1-triazolylmethyl	1-indolylmethyl	2-methylbutyl
2944	1-tetrazolylmethyl	1-triazolylmethyl	2-methylbutyl
2945	2-pyridylmethyl	1-tetrazolylmethyl	2-methylbutyl
2946	3-pyridylmethyl	2-pyridylmethyl	2-methylbutyl
2947	4-pyridylmethyl	3-pyridylmethyl	2-methylbutyl
2948	cyclohexylmethyl	4-pyridylmethyl	2-methylbutyl
2949	2-naphthylmethyl	cyclohexylmethyl	2-methylbutyl
2950	3-naphthylmethyl	2-naphthylmethyl	2-methylbutyl
2951	2-thiophenylmethyl	3-naphthylmethyl	2-methylbutyl
2952	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-methylbutyl
2953	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-methylbutyl
2954	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-methylbutyl
2955	4-biphenylmethyl	2-thienylmethyl	2-methylbutyl
2956	pyrimidinylmethyl	4-biphenylmethyl	2-methylbutyl
2957	2-benzothiazolylmethyl	pyrimidinylmethyl	2-methylbutyl

Table 5

2958	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-methylbutyl
2959	2-thiomethylethyl	2-benzothiophenylmethyl	2-methylbutyl
2960	2-thiomethylmethyl	2-thiomethylethyl	2-methylbutyl
2961	2-methylpropyl	2-thiomethylmethyl	2-methylbutyl
2962	2-methylbutyl	2-methylpropyl	2-methylbutyl
2963	3-methylbutyl	2-methylbutyl	2-methylbutyl
2964	cyclopropylmethyl	3-methylbutyl	2-methylbutyl
2965	cyclobutylmethyl	cyclopropylmethyl	2-methylbutyl
2966	cyclopentylmethyl	cyclobutylmethyl	2-methylbutyl
2967	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-methylbutyl
2968	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-methylbutyl
2969	p-aminophenyl-methyl	p-nitrophenylmethyl	2-methylbutyl
2970	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-methylbutyl
2971	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	propargyl
2972	1-pyrollylmethyl	benzyl	propargyl
2973	1-pyrazolylmethyl	1-pyrollylmethyl	propargyl
2974	1-imidazolylmethyl	1-pyrazolylmethyl	propargyl
2975	1-indolylmethyl	1-imidazolylmethyl	propargyl
2976	1-triazolylmethyl	1-indolylmethyl	propargyl
2977	1-tetrazolylmethyl	1-triazolylmethyl	propargyl
2978	2-pyridylmethyl	1-tetrazolylmethyl	propargyl
2979	3-pyridylmethyl	2-pyridylmethyl	propargyl
2980	4-pyridylmethyl	3-pyridylmethyl	propargyl
2981	cyclohexylmethyl	4-pyridylmethyl	propargyl
2982	2-naphthylmethyl	cyclohexylmethyl	propargyl
2983	3-naphthylmethyl	2-naphthylmethyl	propargyl
2984	2-thiophenylmethyl	3-naphthylmethyl	propargyl
2985	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	propargyl
2986	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	propargyl
2987	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	propargyl
2988	4-biphenylmethyl	2-thienylmethyl	propargyl
2989	pyrimidinylmethyl	4-biphenylmethyl	propargyl
2990	2-benzothiazolylmethyl	pyrimidinylmethyl	propargyl
2991	2-benzothiophenylmethyl	2-benzothiazolylmethyl	propargyl
2992	2-thiomethylethyl	2-benzothiophenylmethyl	propargyl
2993	2-thiomethylmethyl	2-thiomethylethyl	propargyl
2994	2-methylpropyl	2-thiomethylmethyl	propargyl
2995	2-methylbutyl	2-methylpropyl	propargyl
2996	3-methylbutyl	2-methylbutyl	propargyl
2997	cyclopropylmethyl	3-methylbutyl	propargyl
2998	cyclobutylmethyl	cyclopropylmethyl	propargyl
2999	cyclopentylmethyl	cyclobutylmethyl	propargyl
3000	p-hydroxyphenyl-methyl	cyclopentylmethyl	propargyl
3001	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	propargyl

Table 5

3002	p-aminophenyl-methyl	p-nitrophenylmethyl	propargyl	
3003	"4-(N,N-dimethylamino) phenylmethyl"	p-aminophenyl-methyl	propargyl	
3004	benzyl	"4-(N,N-dimethylamino) phenylmethyl"	cyclobutyl	
3005	1-pyrollylmethyl	benzyl	cyclobutyl	
3006	1-pyrazolylmethyl	1-pyrollylmethyl	cyclobutyl	
3007	1-imidazolylmethyl	1-pyrazolylmethyl	cyclobutyl	
3008	1-indolylmethyl	1-imidazolylmethyl	cyclobutyl	
3009	1-triazolylmethyl	1-indolylmethyl	cyclobutyl	
3010	1-tetrazolylmethyl	1-triazolylmethyl	cyclobutyl	
3011	2-pyridylmethyl	1-tetrazolylmethyl	cyclobutyl	
3012	3-pyridylmethyl	2-pyridylmethyl	cyclobutyl	
3013	4-pyridylmethyl	3-pyridylmethyl	cyclobutyl	
3014	cyclohexylmethyl	4-pyridylmethyl	cyclobutyl	
3015	2-naphthylmethyl	cyclohexylmethyl	cyclobutyl	
3016	3-naphthylmethyl	2-naphthylmethyl	cyclobutyl	
3017	2-thiophenylmethyl	3-naphthylmethyl	cyclobutyl	
3018	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	cyclobutyl	
3019	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	cyclobutyl	
3020	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclobutyl	
3021	4-biphenylmethyl	2-thienylmethyl	cyclobutyl	
3022	pyrimidinylmethyl	4-biphenylmethyl	cyclobutyl	
3023	2-benzothiazolylmethyl	pyrimidinylmethyl	cyclobutyl	
3024	2-benzothiophenylmethyl	2-benzothiazolylmethyl	cyclobutyl	
3025	2-thiomethylethyl	2-benzothiophenylmethyl	cyclobutyl	
3026	2-thiomethylmethyl	2-thiomethylethyl	cyclobutyl	
3027	2-methylpropyl	2-thiomethylmethyl	cyclobutyl	
3028	2-methylbutyl	2-methylpropyl	cyclobutyl	
3029	3-methylbutyl	2-methylbutyl	cyclobutyl	
3030	cyclopropylmethyl	3-methylbutyl	cyclobutyl	
3031	cyclobutylmethyl	cyclopropylmethyl	cyclobutyl	
3032	cyclopentylmethyl	cyclobutylmethyl	cyclobutyl	
3033	p-hydroxyphenyl-methyl	cyclopentylmethyl	cyclobutyl	
3034	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	cyclobutyl	
3035	p-aminophenyl-methyl	p-nitrophenylmethyl	cyclobutyl	
3036	"4-(N,N-dimethylamino) phenylmethyl"	p-aminophenyl-methyl	cyclobutyl	
3037	benzyl	"4-(N,N-dimethylamino) phenylmethyl"	cyclobutylmeth yl	
3038	1-pyrollylmethyl	benzyl	cyclobutylmeth yl	
3039	1-pyrazolylmethyl	1-pyrollylmethyl	cyclobutylmeth yl	
3040	1-imidazolylmethyl	1-pyrazolylmethyl	cyclobutylmeth yl	

Table 5

3041	1-indolylmethyl	1-imidazolylmethyl	cyclobutylmeth yl
3042	1-triazolylmethyl	1-indolylmethyl	cyclobutylmeth yl
3043	1-tetrazolylmethyl	1-triazolylmethyl	cyclobutylmeth yl
3044	2-pyridylmethyl	1-tetrazolylmethyl	cyclobutylmeth yl
3045	3-pyridylmethyl	2-pyridylmethyl	cyclobutylmeth yl
3046	4-pyridylmethyl	3-pyridylmethyl	cyclobutylmeth yl
3047	cyclohexylmethyl	4-pyridylmethyl	cyclobutylmeth yl
3048	2-naphthylmethyl	cyclohexylmethyl	cyclobutylmeth yl
3049	3-naphthylmethyl	2-naphthylmethyl	cyclobutylmeth yl
3050	2-thiophenylmethyl	3-naphthylmethyl	cyclobutylmeth yl
3051	4-(1-methyl)piperidinyl- methyl	2-thiophenylmethyl	cyclobutylmeth yl
3052	"(3,4-methylenedioxyphenyl)m ethyl"	4-(1-methyl)piperidinyl- methyl	cyclobutylmeth yl
3053	2-thienylmethyl	"(3,4-methylenedioxyphenyl)m ethyl"	cyclobutylmeth yl
3054	4-biphenylmethyl	2-thienylmethyl	cyclobutylmeth yl
3055	pyrimidinylmethyl	4-biphenylmethyl	cyclobutylmeth yl
3056	2-benzothiazolylmethyl	pyrimidinylmethyl	cyclobutylmeth yl
3057	2-benzothiophenylmethyl	2-benzothiazolylmethyl	cyclobutylmeth yl
3058	2-thiomethylethyl	2-benzothiophenylmethyl	cyclobutylmeth yl
3059	2-thiomethylmethyl	2-thiomethylethyl	cyclobutylmeth yl
3060	2-methylpropyl	2-thiomethylmethyl	cyclobutylmeth yl
3061	2-methylbutyl	2-methylpropyl	cyclobutylmeth yl
3062	3-methylbutyl	2-methylbutyl	cyclobutylmeth yl
3063	cyclopropylmethyl	3-methylbutyl	cyclobutylmeth yl
3064	cyclobutylmethyl	cyclopropylmethyl	cyclobutylmeth yl
3065	cyclopentylmethyl	cyclobutylmethyl	cyclobutylmeth yl
3066	p-hydroxyphenyl-methyl	cyclopentylmethyl	cyclobutylmeth yl
3067	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	cyclobutylmeth yl

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3068	p-aminophenyl-methyl	p-nitrophenylmethyl	cyclobutylmeth yl	+
3069	"4-(N,N- dimethylamino)phenylme thyl"	p-aminophenyl-methyl	cyclobutylmeth yl	2
3070	benzyl	"4-(N,N- dimethylamino)phenylme thyl"	cyclopentyl	4
3071	1-pyrolylmethyl	benzyl	cyclopentyl	2
3072	1-pyrazolylmethyl	1-pyrolylmethyl	cyclopentyl	
3073	1-imidazolylmethyl	1-pyrazolylmethyl	cyclopentyl	
3074	1-indolylmethyl	1-imidazolylmethyl	cyclopentyl	
3075	1-triazolylmethyl	1-indolylmethyl	cyclopentyl	
3076	1-tetrazolylmethyl	1-triazolylmethyl	cyclopentyl	
3077	2-pyridylmethyl	1-tetrazolylmethyl	cyclopentyl	
3078	3-pyridylmethyl	2-pyridylmethyl	cyclopentyl	
3079	4-pyridylmethyl	3-pyridylmethyl	cyclopentyl	
3080	cyclohexylmethyl	4-pyridylmethyl	cyclopentyl	
3081	2-naphthylmethyl	cyclohexylmethyl	cyclopentyl	
3082	3-naphthylmethyl	2-naphthylmethyl	cyclopentyl	
3083	2-thiophenylmethyl	3-naphthylmethyl	cyclopentyl	
3084	4-(1- methyl)piperidinyl- methyl	2-thiophenylmethyl	cyclopentyl	
3085	"(3,4- methylenedioxyphenyl)m ethyl"	4-(1- methyl)piperidinyl- methyl	cyclopentyl	
3086	2-thienylmethyl	"(3,4- methylenedioxyphenyl)m ethyl"	cyclopentyl	
3087	4-biphenylmethyl	2-thienylmethyl	cyclopentyl	
3088	pyrimidinylmethyl	4-biphenylmethyl	cyclopentyl	
3089	2-benzothiazolylmethyl	pyrimidinylmethyl	cyclopentyl	
3090	2- benzothiophenylmethyl	2-benzothiazolylmethyl	cyclopentyl	
3091	2-thiomethylethyl	2- benzothiophenylmethyl	cyclopentyl	
3092	2-thiomethylmethyl	2-thiomethylethyl	cyclopentyl	
3093	2-methylpropyl	2-thiomethylmethyl	cyclopentyl	
3094	2-methylbutyl	2-methylpropyl	cyclopentyl	
3095	3-methylbutyl	2-methylbutyl	cyclopentyl	
3096	cyclopropylmethyl	3-methylbutyl	cyclopentyl	
3097	cyclobutylmethyl	cyclopropylmethyl	cyclopentyl	
3098	cyclopentylmethyl	cyclobutylmethyl	cyclopentyl	
3099	p-hydroxyphenyl-methyl	cyclopentylmethyl	cyclopentyl	
3100	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	cyclopentyl	
3101	p-aminophenyl-methyl	p-nitrophenylmethyl	cyclopentyl	
3102	"4-(N,N- dimethylamino)phenylme thyl"	p-aminophenyl-methyl	cyclopentyl	
3103	benzyl	"4-(N,N- dimethylamino)phenylme thyl"	cyclopentylmet hyl	6
3104	1-pyrolylmethyl	benzyl	cyclopentylmet hyl	2, 7
3105	1-pyrazolylmethyl	1-pyrolylmethyl	cyclopentylmet hyl	8

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3106	1-imidazolylmethyl	1-pyrazolylmethyl	cyclopentylmet hyl
3107	1-indolylmethyl	1-imidazolylmethyl	cyclopentylmet hyl
3108	1-triazolylmethyl	1-indolylmethyl	cyclopentylmet hyl
3109	1-tetrazolylmethyl	1-triazolylmethyl	cyclopentylmet hyl
3110	2-pyridylmethyl	1-tetrazolylmethyl	cyclopentylmet hyl
3111	3-pyridylmethyl	2-pyridylmethyl	cyclopentylmet hyl
3112	4-pyridylmethyl	3-pyridylmethyl	cyclopentylmet hyl
3113	cyclohexylmethyl	4-pyridylmethyl	cyclopentylmet hyl
3114	2-naphthylmethyl	cyclohexylmethyl	cyclopentylmet hyl
3115	3-naphthylmethyl	2-naphthylmethyl	cyclopentylmet hyl
3116	2-thiophenylmethyl	3-naphthylmethyl	cyclopentylmet hyl
3117	4-(1-methyl)piperidinyl-	2-thiophenylmethyl	cyclopentylmet hyl
3118	methyl "(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-	cyclopentylmet hyl
3119	2-thienylmethyl	methyl "(3,4-methylenedioxyphenyl)methyl"	cyclopentylmet hyl
3120	4-biphenylmethyl	2-thienylmethyl	cyclopentylmet hyl
3121	pyrimidinylmethyl	4-biphenylmethyl	cyclopentylmet hyl
3122	2-benzothiazolylmethyl	pyrimidinylmethyl	cyclopentylmet hyl
3123	2-benzothiophenylmethyl	2-benzothiazolylmethyl	cyclopentylmet hyl
3124	2-thiomethylethyl	2-benzothiophenylmethyl	cyclopentylmet hyl
3125	2-thiomethylmethyl	2-thiomethylethyl	cyclopentylmet hyl
3126	2-methylpropyl	2-thiomethylmethyl	cyclopentylmet hyl
3127	2-methylbutyl	2-methylpropyl	cyclopentylmet hyl
3128	3-methylbutyl	2-methylbutyl	cyclopentylmet hyl
3129	cyclopropylmethyl	3-methylbutyl	cyclopentylmet hyl
3130	cyclobutylmethyl	cyclopropylmethyl	cyclopentylmet hyl
3131	cyclopentylmethyl	cyclobutylmethyl	cyclopentylmet hyl
3132	p-hydroxyphenyl-methyl	cyclopentylmethyl	cyclopentylmet hyl

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3133	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	cyclopentylmet hyl
3134	p-aminophenyl-methyl	p-nitrophenylmethyl	cyclopentylmet hyl
3135	"4-(N,N- dimethylamino)phenylme thyl"	p-aminophenyl-methyl	cyclopentylmet hyl
3136	benzyl	"4-(N,N- dimethylamino)phenylme thyl"	1-hexyl
3137	1-pyrolylmethyl	benzyl	1-hexyl
3138	1-pyrazolylmethyl	1-pyrolylmethyl	1-hexyl
3139	1-imidazolylmethyl	1-pyrazolylmethyl	1-hexyl
3140	1-indolylmethyl	1-imidazolylmethyl	1-hexyl
3141	1-triazolylmethyl	1-indolylmethyl	1-hexyl
3142	1-tetrazolylmethyl	1-triazolylmethyl	1-hexyl
3143	2-pyridylmethyl	1-tetrazolylmethyl	1-hexyl
3144	3-pyridylmethyl	2-pyridylmethyl	1-hexyl
3145	4-pyridylmethyl	3-pyridylmethyl	1-hexyl
3146	cyclohexylmethyl	4-pyridylmethyl	1-hexyl
3147	2-naphthylmethyl	cyclohexylmethyl	1-hexyl
3148	3-naphthylmethyl	2-naphthylmethyl	1-hexyl
3149	2-thiophenylmethyl	3-naphthylmethyl	1-hexyl
3150	4-(1- methyl)piperidinyl- methyl	2-thiophenylmethyl	1-hexyl
3151	"(3,4- methylenedioxyphenyl)m ethyl"	4-(1- methyl)piperidinyl- methyl	1-hexyl
3152	2-thienylmethyl	"(3,4- methylenedioxyphenyl)m ethyl"	1-hexyl
3153	4-biphenylmethyl	2-thienylmethyl	1-hexyl
3154	pyrimidinylmethyl	4-biphenylmethyl	1-hexyl
3155	2-benzothiazolylmethyl	pyrimidinylmethyl	1-hexyl
3156	2- benzothiophenylmethyl	2-benzothiazolylmethyl	1-hexyl
3157	2-thiomethylethyl	2- benzothiophenylmethyl	1-hexyl
3158	2-thiomethylmethyl	2-thiomethylethyl	1-hexyl
3159	2-methylpropyl	2-thiomethylmethyl	1-hexyl
3160	2-methylbutyl	2-methylpropyl	1-hexyl
3161	3-methylbutyl	2-methylbutyl	1-hexyl
3162	cyclopropylmethyl	3-methylbutyl	1-hexyl
3163	cyclobutylmethyl	cyclopropylmethyl	1-hexyl
3164	cyclopentylmethyl	cyclobutylmethyl	1-hexyl
3165	p-hydroxyphenyl-methyl	cyclopentylmethyl	1-hexyl
3166	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	1-hexyl
3167	p-aminophenyl-methyl	p-nitrophenylmethyl	1-hexyl
3168	"4-(N,N- dimethylamino)phenylme thyl"	p-aminophenyl-methyl	1-hexyl
3169	benzyl	"4-(N,N- dimethylamino)phenylme thyl"	4-methylpentyl
3170	1-pyrolylmethyl	benzyl	4-methylpentyl
3171	1-pyrazolylmethyl	1-pyrolylmethyl	4-methylpentyl
3172	1-imidazolylmethyl	1-pyrazolylmethyl	4-methylpentyl

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3173	1-indolylmethyl	1-imidazolylmethyl	4-methylpentyl
3174	1-triazolylmethyl	1-indolylmethyl	4-methylpentyl
3175	1-tetrazolylmethyl	1-triazolylmethyl	4-methylpentyl
3176	2-pyridylmethyl	1-tetrazolylmethyl	4-methylpentyl
3177	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl
3178	4-pyridylmethyl	3-pyridylmethyl	4-methylpentyl
3179	cyclohexylmethyl	4-pyridylmethyl	4-methylpentyl
3180	2-naphthylmethyl	cyclohexylmethyl	4-methylpentyl
3181	3-naphthylmethyl	2-naphthylmethyl	4-methylpentyl
3182	2-thiophenylmethyl	3-naphthylmethyl	4-methylpentyl
3183	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	4-methylpentyl
3184	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	4-methylpentyl
3185	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	4-methylpentyl
3186	4-biphenylmethyl	2-thienylmethyl	4-methylpentyl
3187	pyrimidinylmethyl	4-biphenylmethyl	4-methylpentyl
3188	2-benzothiazolylmethyl	pyrimidinylmethyl	4-methylpentyl
3189	2-benzothiophenylmethyl	2-benzothiazolylmethyl	4-methylpentyl
3190	2-thiomethylethyl	2-benzothiophenylmethyl	4-methylpentyl
3191	2-thiomethylmethyl	2-thiomethylethyl	4-methylpentyl
3192	2-methylpropyl	2-thiomethylmethyl	4-methylpentyl
3193	2-methylbutyl	2-methylpropyl	4-methylpentyl
3194	3-methylbutyl	2-methylbutyl	4-methylpentyl
3195	cyclopropylmethyl	3-methylbutyl	4-methylpentyl
3196	cyclobutylmethyl	cyclopropylmethyl	4-methylpentyl
3197	cyclopentylmethyl	cyclobutylmethyl	4-methylpentyl
3198	p-hydroxyphenyl-methyl	cyclopentylmethyl	4-methylpentyl
3199	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	4-methylpentyl
3200	p-aminophenyl-methyl	p-nitrophenylmethyl	4-methylpentyl
3201	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	4-methylpentyl
3202	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	3-methylpentyl
3203	1-pyrollylmethyl	benzyl	3-methylpentyl
3204	1-pyrazolylmethyl	1-pyrollylmethyl	3-methylpentyl
3205	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylpentyl
3206	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl
3207	1-triazolylmethyl	1-indolylmethyl	3-methylpentyl
3208	1-tetrazolylmethyl	1-triazolylmethyl	3-methylpentyl
3209	2-pyridylmethyl	1-tetrazolylmethyl	3-methylpentyl
3210	3-pyridylmethyl	2-pyridylmethyl	3-methylpentyl
3211	4-pyridylmethyl	3-pyridylmethyl	3-methylpentyl
3212	cyclohexylmethyl	4-pyridylmethyl	3-methylpentyl
3213	2-naphthylmethyl	cyclohexylmethyl	3-methylpentyl
3214	3-naphthylmethyl	2-naphthylmethyl	3-methylpentyl
3215	2-thiophenylmethyl	3-naphthylmethyl	3-methylpentyl
3216	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	3-methylpentyl

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3217	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	3-methylpentyl
3218	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-methylpentyl
3219	4-biphenylmethyl	2-thienylmethyl	3-methylpentyl
3220	pyrimidinylmethyl	4-biphenylmethyl	3-methylpentyl
3221	2-benzothiazolylmethyl	pyrimidinylmethyl	3-methylpentyl
3222	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-methylpentyl
3223	2-thiomethylethyl	2-benzothiophenylmethyl	3-methylpentyl
3224	2-thiomethylmethyl	2-thiomethylethyl	3-methylpentyl
3225	2-methylpropyl	2-thiomethylmethyl	3-methylpentyl
3226	2-methylbutyl	2-methylpropyl	3-methylpentyl
3227	3-methylbutyl	2-methylbutyl	3-methylpentyl
3228	cyclopropylmethyl	3-methylbutyl	3-methylpentyl
3229	cyclobutylmethyl	cyclopropylmethyl	3-methylpentyl
3230	cyclopentylmethyl	cyclobutylmethyl	3-methylpentyl
3231	p-hydroxyphenyl-methyl	cyclopentylmethyl	3-methylpentyl
3232	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	3-methylpentyl
3233	p-aminophenyl-methyl	p-nitrophenylmethyl	3-methylpentyl
3234	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	3-methylpentyl
3235	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	isopropyl
3236	1-pyrollylmethyl	benzyl	isopropyl
3237	1-pyrazolylmethyl	1-pyrollylmethyl	isopropyl
3238	1-imidazolylmethyl	1-pyrazolylmethyl	isopropyl
3239	1-indolylmethyl	1-imidazolylmethyl	isopropyl
3240	1-triazolylmethyl	1-indolylmethyl	isopropyl
3241	1-tetrazolylmethyl	1-triazolylmethyl	isopropyl
3242	2-pyridylmethyl	1-tetrazolylmethyl	isopropyl
3243	3-pyridylmethyl	2-pyridylmethyl	isopropyl
3244	4-pyridylmethyl	3-pyridylmethyl	isopropyl
3245	cyclohexylmethyl	4-pyridylmethyl	isopropyl
3246	2-naphthylmethyl	cyclohexylmethyl	isopropyl
3247	3-naphthylmethyl	2-naphthylmethyl	isopropyl
3248	2-thiophenylmethyl	3-naphthylmethyl	isopropyl
3249	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	isopropyl
3250	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	isopropyl
3251	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	isopropyl
3252	4-biphenylmethyl	2-thienylmethyl	isopropyl
3253	pyrimidinylmethyl	4-biphenylmethyl	isopropyl
3254	2-benzothiazolylmethyl	pyrimidinylmethyl	isopropyl
3255	2-benzothiophenylmethyl	2-benzothiazolylmethyl	isopropyl
3256	2-thiomethylethyl	2-benzothiophenylmethyl	isopropyl

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3257	2-thiomethylmethyl	2-thiomethylethyl	isopropyl
3258	2-methylpropyl	2-thiomethylmethyl	isopropyl
3259	2-methylbutyl	2-methylpropyl	isopropyl
3260	3-methylbutyl	2-methylbutyl	isopropyl
3261	cyclopropylmethyl	3-methylbutyl	isopropyl
3262	cyclobutylmethyl	cyclopropylmethyl	isopropyl
3263	cyclopentylmethyl	cyclobutylmethyl	isopropyl
3264	p-hydroxyphenyl-methyl	cyclopentylmethyl	isopropyl
3265	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	isopropyl
3266	p-aminophenyl-methyl	p-nitrophenylmethyl	isopropyl
3267	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	isopropyl
3268	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	3-methylbutyl
3269	1-pyrollylmethyl	benzyl	3-methylbutyl
3270	1-pyrazolylmethyl	1-pyrollylmethyl	3-methylbutyl
3271	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl
3272	1-indolylmethyl	1-imidazolylmethyl	3-methylbutyl
3273	1-triazolylmethyl	1-indolylmethyl	3-methylbutyl
3274	1-tetrazolylmethyl	1-triazolylmethyl	3-methylbutyl
3275	2-pyridylmethyl	1-tetrazolylmethyl	3-methylbutyl
3276	3-pyridylmethyl	2-pyridylmethyl	3-methylbutyl
3277	4-pyridylmethyl	3-pyridylmethyl	3-methylbutyl
3278	cyclohexylmethyl	4-pyridylmethyl	3-methylbutyl
3279	2-naphthylmethyl	cyclohexylmethyl	3-methylbutyl
3280	3-naphthylmethyl	2-naphthylmethyl	3-methylbutyl
3281	2-thiophenylmethyl	3-naphthylmethyl	3-methylbutyl
3282	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	3-methylbutyl
3283	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	3-methylbutyl
3284	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-methylbutyl
3285	4-biphenylmethyl	2-thienylmethyl	3-methylbutyl
3286	pyrimidinylmethyl	4-biphenylmethyl	3-methylbutyl
3287	2-benzothiazolylmethyl	pyrimidinylmethyl	3-methylbutyl
3288	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-methylbutyl
3289	2-thiomethylethyl	2-benzothiophenylmethyl	3-methylbutyl
3290	2-thiomethylmethyl	2-thiomethylethyl	3-methylbutyl
3291	2-methylpropyl	2-thiomethylmethyl	3-methylbutyl
3292	2-methylbutyl	2-methylpropyl	3-methylbutyl
3293	3-methylbutyl	2-methylbutyl	3-methylbutyl
3294	cyclopropylmethyl	3-methylbutyl	3-methylbutyl
3295	cyclobutylmethyl	cyclopropylmethyl	3-methylbutyl
3296	cyclopentylmethyl	cyclobutylmethyl	3-methylbutyl
3297	p-hydroxyphenyl-methyl	cyclopentylmethyl	3-methylbutyl
3298	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	3-methylbutyl
3299	p-aminophenyl-methyl	p-nitrophenylmethyl	3-methylbutyl
3300	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	3-methylbutyl

Table 5

3301	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-phenylethyl
3302	1-pyrolylmethyl	benzyl	2-phenylethyl
3303	1-pyrazolylmethyl	1-pyrolylmethyl	2-phenylethyl
3304	1-imidazolylmethyl	1-pyrazolylmethyl	2-phenylethyl
3305	1-indolylmethyl	1-imidazolylmethyl	2-phenylethyl
3306	1-triazolylmethyl	1-indolylmethyl	2-phenylethyl
3307	1-tetrazolylmethyl	1-triazolylmethyl	2-phenylethyl
3308	2-pyridylmethyl	1-tetrazolylmethyl	2-phenylethyl
3309	3-pyridylmethyl	2-pyridylmethyl	2-phenylethyl
3310	4-pyridylmethyl	3-pyridylmethyl	2-phenylethyl
3311	cyclohexylmethyl	4-pyridylmethyl	2-phenylethyl
3312	2-naphthylmethyl	cyclohexylmethyl	2-phenylethyl
3313	3-naphthylmethyl	2-naphthylmethyl	2-phenylethyl
3314	2-thiophenylmethyl	3-naphthylmethyl	2-phenylethyl
3315	4-(1-methyl)piperidinylmethyl	2-thiophenylmethyl	2-phenylethyl
3316	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinylmethyl	2-phenylethyl
3317	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-phenylethyl
3318	4-biphenylmethyl	2-thienylmethyl	2-phenylethyl
3319	pyrimidinylmethyl	4-biphenylmethyl	2-phenylethyl
3320	2-benzothiazolylmethyl	pyrimidinylmethyl	2-phenylethyl
3321	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-phenylethyl
3322	2-thiomethylethyl	2-benzothiophenylmethyl	2-phenylethyl
3323	2-thiomethylmethyl	2-thiomethylethyl	2-phenylethyl
3324	2-methylpropyl	2-thiomethylmethyl	2-phenylethyl
3325	2-methylbutyl	2-methylpropyl	2-phenylethyl
3326	3-methylbutyl	2-methylbutyl	2-phenylethyl
3327	cyclopropylmethyl	3-methylbutyl	2-phenylethyl
3328	cyclobutylmethyl	cyclopropylmethyl	2-phenylethyl
3329	cyclopentylmethyl	cyclobutylmethyl	2-phenylethyl
3330	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-phenylethyl
3331	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-phenylethyl
3332	p-aminophenyl-methyl	p-nitrophenylmethyl	2-phenylethyl
3333	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-phenylethyl
3334	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	3-phenylpropyl
3335	1-pyrolylmethyl	benzyl	3-phenylpropyl
3336	1-pyrazolylmethyl	1-pyrolylmethyl	3-phenylpropyl
3337	1-imidazolylmethyl	1-pyrazolylmethyl	3-phenylpropyl
3338	1-indolylmethyl	1-imidazolylmethyl	3-phenylpropyl
3339	1-triazolylmethyl	1-indolylmethyl	3-phenylpropyl
3340	1-tetrazolylmethyl	1-triazolylmethyl	3-phenylpropyl
3341	2-pyridylmethyl	1-tetrazolylmethyl	3-phenylpropyl
3342	3-pyridylmethyl	2-pyridylmethyl	3-phenylpropyl
3343	4-pyridylmethyl	3-pyridylmethyl	3-phenylpropyl
3344	cyclohexylmethyl	4-pyridylmethyl	3-phenylpropyl

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3345	2-naphthylmethyl	cyclohexylmethyl	3-phenylpropyl
3346	3-naphthylmethyl	2-naphthylmethyl	3-phenylpropyl
3347	2-thiophenylmethyl	3-naphthylmethyl	3-phenylpropyl
3348	4-(1-methyl)piperidinyl-	2-thiophenylmethyl	3-phenylpropyl
	methyl		
3349	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-	3-phenylpropyl
	ethyl"	methyl	
3350	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-phenylpropyl
3351	4-biphenylmethyl	2-thienylmethyl	3-phenylpropyl
3352	pyrimidinylmethyl	4-biphenylmethyl	3-phenylpropyl
3353	2-benzothiazolylmethyl	pyrimidinylmethyl	3-phenylpropyl
3354	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-phenylpropyl
3355	2-thiomethylethyl	2-benzothiophenylmethyl	3-phenylpropyl
3356	2-thiomethylmethyl	2-thiomethylethyl	3-phenylpropyl
3357	2-methylpropyl	2-thiomethylmethyl	3-phenylpropyl
3358	2-methylbutyl	2-methylpropyl	3-phenylpropyl
3359	3-methylbutyl	2-methylbutyl	3-phenylpropyl
3360	cyclopropylmethyl	3-methylbutyl	3-phenylpropyl
3361	cyclobutylmethyl	cyclopropylmethyl	3-phenylpropyl
3362	cyclopentylmethyl	cyclobutylmethyl	3-phenylpropyl
3363	p-hydroxyphenyl-methyl	cyclopentylmethyl	3-phenylpropyl
3364	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	3-phenylpropyl
3365	p-aminophenyl-methyl	p-nitrophenylmethyl	3-phenylpropyl
3366	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	3-phenylpropyl
	benzyl		
3367		"4-(N,N-dimethylamino)phenylmethyl"	"2-(N,N-dimethylamino)ethyl"
		benzyl	"2-(N,N-dimethylamino)ethyl"
3368	1-pyrollylmethyl		"2-(N,N-dimethylamino)ethyl"
			"2-(N,N-dimethylamino)ethyl"
3369	1-pyrazolylmethyl	1-pyrollylmethyl	"2-(N,N-dimethylamino)ethyl"
			"2-(N,N-dimethylamino)ethyl"
3370	1-imidazolylmethyl	1-pyrazolylmethyl	"2-(N,N-dimethylamino)ethyl"
			"2-(N,N-dimethylamino)ethyl"
3371	1-indolylmethyl	1-imidazolylmethyl	"2-(N,N-dimethylamino)ethyl"
			"2-(N,N-dimethylamino)ethyl"
3372	1-triazolylmethyl	1-indolylmethyl	"2-(N,N-dimethylamino)ethyl"
			"2-(N,N-dimethylamino)ethyl"
3373	1-tetrazolylmethyl	1-triazolylmethyl	"2-(N,N-dimethylamino)ethyl"
			"2-(N,N-dimethylamino)ethyl"
3374	2-pyridylmethyl	1-tetrazolylmethyl	"2-(N,N-dimethylamino)ethyl"

Table 5

3375	3-pyridylmethyl	2-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
3376	4-pyridylmethyl	3-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
3377	cyclohexylmethyl	4-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"
3378	2-naphthylmethyl	cyclohexylmethyl	"2-(N,N-dimethylamino)ethyl"
3379	3-naphthylmethyl	2-naphthylmethyl	"2-(N,N-dimethylamino)ethyl"
3380	2-thiophenylmethyl	3-naphthylmethyl	"2-(N,N-dimethylamino)ethyl"
3381	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	"2-(N,N-dimethylamino)ethyl"
3382	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	"2-(N,N-dimethylamino)ethyl"
3383	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	"2-(N,N-dimethylamino)ethyl"
3384	4-biphenylmethyl	2-thienylmethyl	"2-(N,N-dimethylamino)ethyl"
3385	pyrimidinylmethyl	4-biphenylmethyl	"2-(N,N-dimethylamino)ethyl"
3386	2-benzothiazolylmethyl	pyrimidinylmethyl	"2-(N,N-dimethylamino)ethyl"
3387	2-benzothiophenylmethyl	2-benzothiazolylmethyl	"2-(N,N-dimethylamino)ethyl"
3388	2-thiomethylethyl	2-benzothiophenylmethyl	"2-(N,N-dimethylamino)ethyl"
3389	2-thiomethylmethyl	2-thiomethylethyl	"2-(N,N-dimethylamino)ethyl"
3390	2-methylpropyl	2-thiomethylmethyl	"2-(N,N-dimethylamino)ethyl"
3391	2-methylbutyl	2-methylpropyl	"2-(N,N-dimethylamino)ethyl"
3392	3-methylbutyl	2-methylbutyl	"2-(N,N-dimethylamino)ethyl"
3393	cyclopropylmethyl	3-methylbutyl	"2-(N,N-dimethylamino)ethyl"

Table 5

3394	cyclobutylmethyl	cyclopropylmethyl	"2-(N,N-dimethylamino)ethyl"
3395	cyclopentylmethyl	cyclobutylmethyl	"2-(N,N-dimethylamino)ethyl"
3396	p-hydroxyphenyl-methyl	cyclopentylmethyl	"2-(N,N-dimethylamino)ethyl"
3397	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	"2-(N,N-dimethylamino)ethyl"
3398	p-aminophenyl-methyl	p-nitrophenylmethyl	"2-(N,N-dimethylamino)ethyl"
3399	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	"2-(N,N-dimethylamino)ethyl"
3400	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	3-oxetanylmethyl
3401	1-pyrollylmethyl	benzyl	3-oxetanylmethyl
3402	1-pyrazolylmethyl	1-pyrollylmethyl	3-oxetanylmethyl
3403	1-imidazolylmethyl	1-pyrazolylmethyl	3-oxetanylmethyl
3404	1-indolylmethyl	1-imidazolylmethyl	3-oxetanylmethyl
3405	1-triazolylmethyl	1-indolylmethyl	3-oxetanylmethyl
3406	1-tetrazolylmethyl	1-triazolylmethyl	3-oxetanylmethyl
3407	2-pyridylmethyl	1-tetrazolylmethyl	3-oxetanylmethyl
3408	3-pyridylmethyl	2-pyridylmethyl	3-oxetanylmethyl
3409	4-pyridylmethyl	3-pyridylmethyl	3-oxetanylmethyl
3410	cyclohexylmethyl	4-pyridylmethyl	3-oxetanylmethyl
3411	2-naphthylmethyl	cyclohexylmethyl	3-oxetanylmethyl
3412	3-naphthylmethyl	2-naphthylmethyl	3-oxetanylmethyl
3413	2-thiophenylmethyl	3-naphthylmethyl	3-oxetanylmethyl
3414	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	3-oxetanylmethyl
3415	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	3-oxetanylmethyl
3416	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-oxetanylmethyl
3417	4-biphenylmethyl	2-thienylmethyl	3-oxetanylmethyl

Table 5

3418	pyrimidinylmethyl	4-biphenylmethyl	3-oxetanylmethyl	*
3419	2-benzothiazolylmethyl	pyrimidinylmethyl	3-oxetanylmethyl	*
3420	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-oxetanylmethyl	*
3421	2-thiomethylethyl	2-benzothiophenylmethyl	3-oxetanylmethyl	*
3422	2-thiomethylmethyl	2-thiomethylethyl	3-oxetanylmethyl	
3423	2-methylpropyl	2-thiomethylmethyl	3-oxetanylmethyl	
3424	2-methylbutyl	2-methylpropyl	3-oxetanylmethyl	
3425	3-methylbutyl	2-methylbutyl	3-oxetanylmethyl	
3426	cyclopropylmethyl	3-methylbutyl	3-oxetanylmethyl	
3427	cyclobutylmethyl	cyclopropylmethyl	3-oxetanylmethyl	
3428	cyclopentylmethyl	cyclobutylmethyl	3-oxetanylmethyl	
3429	p-hydroxyphenyl-methyl	cyclopentylmethyl	3-oxetanylmethyl	
3430	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	3-oxetanylmethyl	
3431	p-aminophenyl-methyl	p-nitrophenylmethyl	3-oxetanylmethyl	
3432	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	3-oxetanylmethyl	
3433	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-tetrahydrofuran	
3434	1-pyrollylmethyl	benzyl	2-tetrahydrofuran	
3435	1-pyrazolylmethyl	1-pyrollylmethyl	2-tetrahydrofuran	
3436	1-imidazolylmethyl	1-pyrazolylmethyl	2-tetrahydrofuran	
3437	1-indolylmethyl	1-imidazolylmethyl	2-tetrahydrofuran	
3438	1-triazolylmethyl	1-indolylmethyl	2-tetrahydrofuran	
3439	1-tetrazolylmethyl	1-triazolylmethyl	2-tetrahydrofuran	
3440	2-pyridylmethyl	1-tetrazolylmethyl	2-tetrahydrofuran	
3441	3-pyridylmethyl	2-pyridylmethyl	2-tetrahydrofuran	

Table 5

3442	4-pyridylmethyl	3-pyridylmethyl	2-tetrahydrofuran
			nyl
3443	cyclohexylmethyl	4-pyridylmethyl	2-tetrahydrofuran
			nyl
3444	2-naphthylmethyl	cyclohexylmethyl	2-tetrahydrofuran
			nyl
3445	3-naphthylmethyl	2-naphthylmethyl	2-tetrahydrofuran
			nyl
3446	2-thiophenylmethyl	3-naphthylmethyl	2-tetrahydrofuran
			nyl
3447	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-tetrahydrofuran
			nyl
3448	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-tetrahydrofuran
			nyl
3449	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-tetrahydrofuran
			nyl
3450	4-biphenylmethyl	2-thienylmethyl	2-tetrahydrofuran
			nyl
3451	pyrimidinylmethyl	4-biphenylmethyl	2-tetrahydrofuran
			nyl
3452	2-benzothiazolylmethyl	pyrimidinylmethyl	2-tetrahydrofuran
			nyl
3453	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-tetrahydrofuran
			nyl
3454	2-thiomethylethyl	2-benzothiophenylmethyl	2-tetrahydrofuran
			nyl
3455	2-thiomethylmethyl	2-thiomethylethyl	2-tetrahydrofuran
			nyl
3456	2-methylpropyl	2-thiomethylmethyl	2-tetrahydrofuran
			nyl
3457	2-methylbutyl	2-methylpropyl	2-tetrahydrofuran
			nyl
3458	3-methylbutyl	2-methylbutyl	2-tetrahydrofuran
			nyl
3459	cyclopropylmethyl	3-methylbutyl	2-tetrahydrofuran
			nyl
3460	cyclobutylmethyl	cyclopropylmethyl	2-tetrahydrofuran
			nyl

Table 5

3461	cyclopentylmethyl	cyclobutylmethyl	2-tetrahydrofuran
3462	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-tetrahydrofuran
3463	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-tetrahydrofuran
3464	p-aminophenyl-methyl	p-nitrophenylmethyl	2-tetrahydrofuran
3465	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-tetrahydrofuran
3466	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-methoxypropyl
3467	1-pyrollylmethyl	benzyl	2-methoxypropyl
3468	1-pyrazolylmethyl	1-pyrollylmethyl	2-methoxypropyl
3469	1-imidazolylmethyl	1-pyrazolylmethyl	2-methoxypropyl
3470	1-indolylmethyl	1-imidazolylmethyl	2-methoxypropyl
3471	1-triazolylmethyl	1-indolylmethyl	2-methoxypropyl
3472	1-tetrazolylmethyl	1-triazolylmethyl	2-methoxypropyl
3473	2-pyridylmethyl	1-tetrazolylmethyl	2-methoxypropyl
3474	3-pyridylmethyl	2-pyridylmethyl	2-methoxypropyl
3475	4-pyridylmethyl	3-pyridylmethyl	2-methoxypropyl
3476	cyclohexylmethyl	4-pyridylmethyl	2-methoxypropyl
3477	2-naphthylmethyl	cyclohexylmethyl	2-methoxypropyl
3478	3-naphthylmethyl	2-naphthylmethyl	2-methoxypropyl
3479	2-thiophenylmethyl	3-naphthylmethyl	2-methoxypropyl
3480	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-methoxypropyl
3481	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-methoxypropyl
3482	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-methoxypropyl
3483	4-biphenylmethyl	2-thienylmethyl	2-methoxypropyl
3484	pyrimidinylmethyl	4-biphenylmethyl	2-methoxypropyl

Table 5

3485	2-benzothiazolylmethyl	pyrimidinylmethyl	2-methoxypropyl
3486	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-methoxypropyl
3487	2-thiomethylethyl	2-benzothiophenylmethyl	2-methoxypropyl
3488	2-thiomethylmethyl	2-thiomethylethyl	2-methoxypropyl
3489	2-methylpropyl	2-thiomethylmethyl	2-methoxypropyl
3490	2-methylbutyl	2-methylpropyl	2-methoxypropyl
3491	3-methylbutyl	2-methylbutyl	2-methoxypropyl
3492	cyclopropylmethyl	3-methylbutyl	2-methoxypropyl
3493	cyclobutylmethyl	cyclopropylmethyl	2-methoxypropyl
3494	cyclopentylmethyl	cyclobutylmethyl	2-methoxypropyl
3495	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-methoxypropyl
3496	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-methoxypropyl
3497	p-aminophenyl-methyl	p-nitrophenylmethyl	2-methoxypropyl
3498	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-methoxypropyl
3499	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-ethoxyethyl
3500	1-pyrollylmethyl	benzyl	2-ethoxyethyl
3501	1-pyrazolylmethyl	1-pyrollylmethyl	2-ethoxyethyl
3502	1-imidazolylmethyl	1-pyrazolylmethyl	2-ethoxyethyl
3503	1-indolylmethyl	1-imidazolylmethyl	2-ethoxyethyl
3504	1-triazolylmethyl	1-indolylmethyl	2-ethoxyethyl
3505	1-tetrazolylmethyl	1-triazolylmethyl	2-ethoxyethyl
3506	2-pyridylmethyl	1-tetrazolylmethyl	2-ethoxyethyl
3507	3-pyridylmethyl	2-pyridylmethyl	2-ethoxyethyl
3508	4-pyridylmethyl	3-pyridylmethyl	2-ethoxyethyl
3509	cyclohexylmethyl	4-pyridylmethyl	2-ethoxyethyl
3510	2-naphthylmethyl	cyclohexylmethyl	2-ethoxyethyl
3511	3-naphthylmethyl	2-naphthylmethyl	2-ethoxyethyl
3512	2-thiophenylmethyl	3-naphthylmethyl	2-ethoxyethyl
3513	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-ethoxyethyl
3514	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-ethoxyethyl
3515	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-ethoxyethyl
3516	4-biphenylmethyl	2-thienylmethyl	2-ethoxyethyl
3517	pyrimidinylmethyl	4-biphenylmethyl	2-ethoxyethyl
3518	2-benzothiazolylmethyl	pyrimidinylmethyl	2-ethoxyethyl

Table 5

3519	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-ethoxyethyl
3520	2-thiomethylethyl	2-benzothiophenylmethyl	2-ethoxyethyl
3521	2-thiomethylmethyl	2-thiomethylethyl	2-ethoxyethyl
3522	2-methylpropyl	2-thiomethylmethyl	2-ethoxyethyl
3523	2-methylbutyl	2-methylpropyl	2-ethoxyethyl
3524	3-methylbutyl	2-methylbutyl	2-ethoxyethyl
3525	cyclopropylmethyl	3-methylbutyl	2-ethoxyethyl
3526	cyclobutylmethyl	cyclopropylmethyl	2-ethoxyethyl
3527	cyclopentylmethyl	cyclobutylmethyl	2-ethoxyethyl
3528	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-ethoxyethyl
3529	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-ethoxyethyl
3530	p-aminophenyl-methyl	p-nitrophenylmethyl	2-ethoxyethyl
3531	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-ethoxyethyl
3532	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-(1-pyrolyl)ethyl
3533	1-pyrolylmethyl	benzyl	2-(1-pyrolyl)ethyl
3534	1-pyrazolylmethyl	1-pyrolylmethyl	2-(1-pyrolyl)ethyl
3535	1-imidazolylmethyl	1-pyrazolylmethyl	2-(1-pyrolyl)ethyl
3536	1-indolylmethyl	1-imidazolylmethyl	2-(1-pyrolyl)ethyl
3537	1-triazolylmethyl	1-indolylmethyl	2-(1-pyrolyl)ethyl
3538	1-tetrazolylmethyl	1-triazolylmethyl	2-(1-pyrolyl)ethyl
3539	2-pyridylmethyl	1-tetrazolylmethyl	2-(1-pyrolyl)ethyl
3540	3-pyridylmethyl	2-pyridylmethyl	2-(1-pyrolyl)ethyl
3541	4-pyridylmethyl	3-pyridylmethyl	2-(1-pyrolyl)ethyl
3542	cyclohexylmethyl	4-pyridylmethyl	2-(1-pyrolyl)ethyl
3543	2-naphthylmethyl	cyclohexylmethyl	2-(1-pyrolyl)ethyl
3544	3-naphthylmethyl	2-naphthylmethyl	2-(1-pyrolyl)ethyl
3545	2-thiophenylmethyl	3-naphthylmethyl	2-(1-pyrolyl)ethyl
3546	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-(1-pyrolyl)ethyl
3547	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-(1-pyrolyl)ethyl
3548	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-(1-pyrolyl)ethyl
3549	4-biphenylmethyl	2-thienylmethyl	2-(1-pyrolyl)ethyl

Table 5

3550	pyrimidinylmethyl	4-biphenylmethyl	2-(1-pyrolyl)ethyl
3551	2-benzothiazolylmethyl	pyrimidinylmethyl	2-(1-pyrolyl)ethyl
3552	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-(1-pyrolyl)ethyl
3553	2-thiomethylethyl	2-benzothiophenylmethyl	2-(1-pyrolyl)ethyl
3554	2-thiomethylmethyl	2-thiomethylethyl	2-(1-pyrolyl)ethyl
3555	2-methylpropyl	2-thiomethylmethyl	2-(1-pyrolyl)ethyl
3556	2-methylbutyl	2-methylpropyl	2-(1-pyrolyl)ethyl
3557	3-methylbutyl	2-methylbutyl	2-(1-pyrolyl)ethyl
3558	cyclopropylmethyl	3-methylbutyl	2-(1-pyrolyl)ethyl
3559	cyclobutylmethyl	cyclopropylmethyl	2-(1-pyrolyl)ethyl
3560	cyclopentylmethyl	cyclobutylmethyl	2-(1-pyrolyl)ethyl
3561	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-(1-pyrolyl)ethyl
3562	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-(1-pyrolyl)ethyl
3563	p-aminophenyl-methyl	p-nitrophenylmethyl	2-(1-pyrolyl)ethyl
3564	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-(1-pyrolyl)ethyl
3565	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-(1-imidazolyl)ethyl
3566	1-pyrollylmethyl	benzyl	2-(1-imidazolyl)ethyl
3567	1-pyrazolylmethyl	1-pyrollylmethyl	2-(1-imidazolyl)ethyl
3568	1-imidazolylmethyl	1-pyrazolylmethyl	2-(1-imidazolyl)ethyl
3569	1-indolylmethyl	1-imidazolylmethyl	2-(1-imidazolyl)ethyl
3570	1-triazolylmethyl	1-indolylmethyl	2-(1-imidazolyl)ethyl
3571	1-tetrazolylmethyl	1-triazolylmethyl	2-(1-imidazolyl)ethyl
3572	2-pyridylmethyl	1-tetrazolylmethyl	2-(1-imidazolyl)ethyl
3573	3-pyridylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl

Table 5

3574	4-pyridylmethyl	3-pyridylmethyl	2-(1-imidazolyl)ethyl	
3575	cyclohexylmethyl	4-pyridylmethyl	2-(1-imidazolyl)ethyl	
3576	2-naphthylmethyl	cyclohexylmethyl	2-(1-imidazolyl)ethyl	
3577	3-naphthylmethyl	2-naphthylmethyl	2-(1-imidazolyl)ethyl	
3578	2-thiophenylmethyl	3-naphthylmethyl	2-(1-imidazolyl)ethyl	
3579	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-(1-imidazolyl)ethyl	
3580	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-(1-imidazolyl)ethyl	
3581	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-(1-imidazolyl)ethyl	
3582	4-biphenylmethyl	2-thienylmethyl	2-(1-imidazolyl)ethyl	
3583	pyrimidinylmethyl	4-biphenylmethyl	2-(1-imidazolyl)ethyl	
3584	2-benzothiazolylmethyl	pyrimidinylmethyl	2-(1-imidazolyl)ethyl	
3585	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-(1-imidazolyl)ethyl	
3586	2-thiomethylethyl	2-benzothiophenylmethyl	2-(1-imidazolyl)ethyl	
3587	2-thiomethylmethyl	2-thiomethylethyl	2-(1-imidazolyl)ethyl	
3588	2-methylpropyl	2-thiomethylmethyl	2-(1-imidazolyl)ethyl	
3589	2-methylbutyl	2-methylpropyl	2-(1-imidazolyl)ethyl	
3590	3-methylbutyl	2-methylbutyl	2-(1-imidazolyl)ethyl	
3591	cyclopropylmethyl	3-methylbutyl	2-(1-imidazolyl)ethyl	
3592	cyclobutylmethyl	cyclopropylmethyl	2-(1-imidazolyl)ethyl	

Table 5

3593	cyclopentylmethyl	cyclobutylmethyl	2-(1-imidazolyl)ethyl
3594	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-(1-imidazolyl)ethyl
3595	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-(1-imidazolyl)ethyl
3596	p-aminophenyl-methyl	p-nitrophenylmethyl	2-(1-imidazolyl)ethyl
3597	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-(1-imidazolyl)ethyl
3598	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-pyridylmethyl
3599	1-pyrollylmethyl	benzyl	2-pyridylmethyl
3600	1-pyrazolylmethyl	1-pyrollylmethyl	2-pyridylmethyl
3601	1-imidazolylmethyl	1-pyrazolylmethyl	2-pyridylmethyl
3602	1-indolylmethyl	1-imidazolylmethyl	2-pyridylmethyl
3603	1-triazolylmethyl	1-indolylmethyl	2-pyridylmethyl
3604	1-tetrazolylmethyl	1-triazolylmethyl	2-pyridylmethyl
3605	2-pyridylmethyl	1-tetrazolylmethyl	2-pyridylmethyl
3606	3-pyridylmethyl	2-pyridylmethyl	2-pyridylmethyl
3607	4-pyridylmethyl	3-pyridylmethyl	2-pyridylmethyl
3608	cyclohexylmethyl	4-pyridylmethyl	2-pyridylmethyl
3609	2-naphthylmethyl	cyclohexylmethyl	2-pyridylmethyl
3610	3-naphthylmethyl	2-naphthylmethyl	2-pyridylmethyl
3611	2-thiophenylmethyl	3-naphthylmethyl	2-pyridylmethyl
3612	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-pyridylmethyl
3613	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-pyridylmethyl
3614	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-pyridylmethyl
3615	4-biphenylmethyl	2-thienylmethyl	2-pyridylmethyl
3616	pyrimidinylmethyl	4-biphenylmethyl	2-pyridylmethyl

Table 5

3617	2-benzothiazolylmethyl	pyrimidinylmethyl	2-pyridylmethyl
3618	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-pyridylmethyl
3619	2-thiomethylethyl	2-benzothiophenylmethyl	2-pyridylmethyl
3620	2-thiomethylmethyl	2-thiomethylethyl	2-pyridylmethyl
3621	2-methylpropyl	2-thiomethylmethyl	2-pyridylmethyl
3622	2-methylbutyl	2-methylpropyl	2-pyridylmethyl
3623	3-methylbutyl	2-methylbutyl	2-pyridylmethyl
3624	cyclopropylmethyl	3-methylbutyl	2-pyridylmethyl
3625	cyclobutylmethyl	cyclopropylmethyl	2-pyridylmethyl
3626	cyclopentylmethyl	cyclobutylmethyl	2-pyridylmethyl
3627	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-pyridylmethyl
3628	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-pyridylmethyl
3629	p-aminophenyl-methyl	p-nitrophenylmethyl	2-pyridylmethyl
3630	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-pyridylmethyl
3631	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-thiomethylethyl
3632	1-pyrollylmethyl	benzyl	1-thiomethylethyl
3633	1-pyrazolylmethyl	1-pyrollylmethyl	1-thiomethylethyl
3634	1-imidazolylmethyl	1-pyrazolylmethyl	1-thiomethylethyl
3635	1-indolylmethyl	1-imidazolylmethyl	1-thiomethylethyl
3636	1-triazolylmethyl	1-indolylmethyl	1-thiomethylethyl
3637	1-tetrazolylmethyl	1-triazolylmethyl	1-thiomethylethyl
3638	2-pyridylmethyl	1-tetrazolylmethyl	1-thiomethylethyl
3639	3-pyridylmethyl	2-pyridylmethyl	1-thiomethylethyl

Table 5

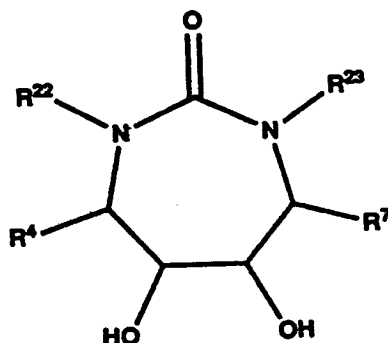
3640	4-pyridylmethyl	3-pyridylmethyl	2-thiomethylethy 1
3641	cyclohexylmethyl	4-pyridylmethyl	2-thiomethylethy 1
3642	2-naphthylmethyl	cyclohexylmethyl	2-thiomethylethy 1
3643	3-naphthylmethyl	2-naphthylmethyl	2-thiomethylethy 1
3644	2-thiophenylmethyl	3-naphthylmethyl	2-thiomethylethy 1
3645	4-(1-methyl)piperidinyl- methyl	2-thiophenylmethyl	2-thiomethylethy 1
3646	"(3,4-methylenedioxyphenyl)m ethyl"	4-(1-methyl)piperidinyl- methyl	2-thiomethylethy 1
3647	2-thienylmethyl	"(3,4-methylenedioxyphenyl)m ethyl"	2-thiomethylethy 1
3648	4-biphenylmethyl	2-thienylmethyl	2-thiomethylethy 1
3649	pyrimidinylmethyl	4-biphenylmethyl	2-thiomethylethy 1
3650	2-benzothiazolylmethyl	pyrimidinylmethyl	2-thiomethylethy 1
3651	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-thiomethylethy 1
3652	2-thiomethylethyl	2-benzothiophenylmethyl	2-thiomethylethy 1
3653	2-thiomethylmethyl	2-thiomethylethyl	2-thiomethylethy 1
3654	2-methylpropyl	2-thiomethylmethyl	2-thiomethylethy 1
3655	2-methylbutyl	2-methylpropyl	2-thiomethylethy 1
3656	3-methylbutyl	2-methylbutyl	2-thiomethylethy 1
3657	cyclopropylmethyl	3-methylbutyl	2-thiomethylethy 1
3658	cyclobutylmethyl	cyclopropylmethyl	2-thiomethylethy 1

Table 5

3659	cyclopentylmethyl	cyclobutylmethyl	2- thiomethylethy 1	
3660	p-hydroxyphenyl-methyl	cyclopentylmethyl	2- thiomethylethy 1	
3661	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2- thiomethylethy 1	
3662	p-aminophenyl-methyl	p-nitrophenylmethyl	2- thiomethylethy 1	
3663	"4-(N,N- dimethylamino)phenylme thyl"	p-aminophenyl-methyl	2- thiomethylethy 1	
3664		"4-(N,N- dimethylamino)phenylme thyl"		

Table 6

TABLE 6

R⁴; R⁷; R²²; R²³

Example Number	R ⁴	R ⁷	R ²²	R ²³
3665	benzyl	"4-(N,N-di-methyl-amino)-phenylmethyl"	allyl	2-thiomethylethyl
3666	1-pyrolylmethyl	benzyl	allyl	2-thiomethylethyl
3667	1-pyrazolylmethyl	1-pyrolylmethyl	allyl	2-thiomethylethyl
3668	1-imidazolylmethyl	1-pyrazolylmethyl	allyl	2-thiomethylethyl
3669	1-indolylmethyl	1-imidazolylmethyl	allyl	2-thiomethylethyl
3670	1-triazolylmethyl	1-indolylmethyl	allyl	2-thiomethylethyl
3671	1-tetrazolylmethyl	1-triazolylmethyl	allyl	2-thiomethylethyl
3672	2-pyridylmethyl	1-tetrazolylmethyl	allyl	2-thiomethylethyl
3673	3-pyridylmethyl	2-pyridylmethyl	allyl	2-thiomethylethyl
3674	4-pyridylmethyl	3-pyridylmethyl	allyl	2-thiomethylethyl
3675	cyclohexylmethyl	4-pyridylmethyl	allyl	2-thiomethylethyl

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3676	2-naphthylmethyl	cyclohexylmethyl	allyl	2-thiomethylethyl
3677	3-naphthylmethyl	2-naphthylmethyl	allyl	2-thiomethylethyl
3678	2-thiophenylmethyl	3-naphthylmethyl	allyl	2-thiomethylethyl
3679	4-(1-methyl)piperidinylmethyl	2-thiophenylmethyl	allyl	2-thiomethylethyl
3680	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	allyl	2-thiomethylethyl
3681	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	allyl	2-thiomethylethyl
3682	4-biphenylmethyl	2-thienylmethyl	allyl	2-thiomethylethyl
3683	pyrimidinylmethyl	4-biphenylmethyl	allyl	2-thiomethylethyl
3684	2-benzothiazolyl-methyl	pyrimidinylmethyl	allyl	2-thiomethylethyl
3685	2-benzothiophenyl-methyl	2-benzothiazolyl-methyl	allyl	2-thiomethylethyl
3686	2-thiomethylethyl	2-benzothiophenyl-methyl	allyl	2-thiomethylethyl
3687	2-thiomethylmethyl	2-thiomethylethyl	allyl	2-thiomethylethyl
3688	2-methylpropyl	2-thiomethylmethyl	allyl	2-thiomethylethyl
3689	2-methylbutyl	2-methylpropyl	allyl	2-thiomethylethyl
3690	3-methylbutyl	2-methylbutyl	allyl	2-thiomethylethyl
3691	cyclopropylmethyl	3-methylbutyl	allyl	2-thiomethylethyl
3692	cyclobutylmethyl	cyclopropylmethyl	allyl	2-thiomethylethyl
3693	cyclopentylmethyl	cyclobutylmethyl	allyl	2-thiomethylethyl
3694	p-hydroxyphenyl-methyl	cyclopentylmethyl	allyl	2-thiomethylethyl
3695	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	allyl	2-thiomethylethyl
3696	p-aminophenyl-methyl	p-nitrophenylmethyl	allyl	2-thiomethylethyl

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3697	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	allyl	2-thiomethylethyl
3698	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	propyl	allyl
3699	1-pyrolylmethyl	benzyl	propyl	allyl
3700	1-pyrazolylmethyl	1-pyrolylmethyl	propyl	allyl
3701	1-imidazolylmethyl	1-pyrazolylmethyl	propyl	allyl
3702	1-indolylmethyl	1-imidazolylmethyl	propyl	allyl
3703	1-triazolylmethyl	1-indolylmethyl	propyl	allyl
3704	1-tetrazolylmethyl	1-triazolylmethyl	propyl	allyl
3705	2-pyridylmethyl	1-tetrazolylmethyl	propyl	allyl
3706	3-pyridylmethyl	2-pyridylmethyl	propyl	allyl
3707	4-pyridylmethyl	3-pyridylmethyl	propyl	allyl
3708	cyclohexylmethyl	4-pyridylmethyl	propyl	allyl
3709	2-naphthylmethyl	cyclohexylmethyl	propyl	allyl
3710	3-naphthylmethyl	2-naphthylmethyl	propyl	allyl
3711	2-thiophenylmethyl	3-naphthylmethyl	propyl	allyl
3712	4-(1-methyl)piperidinylmethyl	2-thiophenylmethyl	propyl	allyl
3713	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinylmethyl	propyl	allyl
3714	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	propyl	allyl
3715	4-biphenylmethyl	2-thienylmethyl	propyl	allyl
3716	pyrimidinylmethyl	4-biphenylmethyl	propyl	allyl
3717	2-benzothiazolylmethyl	pyrimidinylmethyl	propyl	allyl
3718	2-benzothiophenylmethyl	2-benzothiazolylmethyl	propyl	allyl

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3719	2-thiomethylethyl	2-benzothiophenylmethyl	propyl	allyl
3720	2-thiomethylmethyl	2-thiomethylethyl	propyl	allyl
3721	2-methylpropyl	2-thiomethylmethyl	propyl	allyl
3722	2-methylbutyl	2-methylpropyl	propyl	allyl
3723	3-methylbutyl	2-methylbutyl	propyl	allyl
3724	cyclopropylmethyl	3-methylbutyl	propyl	allyl
3725	cyclobutylmethyl	cyclopropylmethyl	propyl	allyl
3726	cyclopentylmethyl	cyclobutylmethyl	propyl	allyl
3727	p-hydroxyphenylmethyl	cyclopentylmethyl	propyl	allyl
3728	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	allyl
3729	p-aminophenylmethyl	p-nitrophenylmethyl	propyl	allyl
3730	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	propyl	allyl
3731	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	n-butyl	propyl
3732	1-pyrollylmethyl	benzyl	n-butyl	propyl
3733	1-pyrazolylmethyl	1-pyrollylmethyl	n-butyl	propyl
3734	1-imidazolylmethyl	1-pyrazolylmethyl	n-butyl	propyl
3735	1-indolylmethyl	1-imidazolylmethyl	n-butyl	propyl
3736	1-triazolylmethyl	1-indolylmethyl	n-butyl	propyl
3737	1-tetrazolylmethyl	1-triazolylmethyl	n-butyl	propyl
3738	2-pyridylmethyl	1-tetrazolylmethyl	n-butyl	propyl
3739	3-pyridylmethyl	2-pyridylmethyl	n-butyl	propyl
3740	4-pyridylmethyl	3-pyridylmethyl	n-butyl	propyl
3741	cyclohexylmethyl	4-pyridylmethyl	n-butyl	propyl
3742	2-naphthylmethyl	cyclohexylmethyl	n-butyl	propyl

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3743	3-naphthylmethyl	2-naphthylmethyl	n-butyl	propyl
3744	2-thiophenylmeth yl	3-naphthylmethyl	n-butyl	propyl
3745	4-(1-methyl)piperid inyl-methyl	2-thiophenylmethyl	n-butyl	propyl
3746	"(3,4-methylenedioxy phenyl)methyl"	4-(1-methyl)piperidin yl-methyl	n-butyl	propyl
3747	2-thienylmethyl	"(3,4-methylenedioxyph enyl)methyl"	n-butyl	propyl
3748	4-biphenylmethyl	2-thienylmethyl	n-butyl	propyl
3749	pyrimidinylmet hyl	4-biphenylmethyl	n-butyl	propyl
3750	2-benzothiazolyl methyl	pyrimidinylmethy l	n-butyl	propyl
3751	2-benzothiopheny lmethyl	2-benzothiazolylme thyl	n-butyl	propyl
3752	2-thiomethylethy l	2-benzothiophenylm ethyl	n-butyl	propyl
3753	2-thiomethylmeth yl	2-thiomethylethyl	n-butyl	propyl
3754	2-methylpropyl	2-thiomethylmethyl	n-butyl	propyl
3755	2-methylbutyl	2-methylpropyl	n-butyl	propyl
3756	3-methylbutyl	2-methylbutyl	n-butyl	propyl
3757	cyclopropylmet hyl	3-methylbutyl	n-butyl	propyl
3758	cyclobutylmeth yl	cyclopropylmethy l	n-butyl	propyl
3759	cyclopentylmet hyl	cyclobutylmethyl	n-butyl	propyl
3760	p-hydroxyphenyl- methyl	cyclopentylmethy l	n-butyl	propyl
3761	p-nitrophenylmet hyl	p-hydroxyphenyl- methyl	n-butyl	propyl
3762	p-aminophenyl- methyl	p-nitrophenylmethy l	n-butyl	propyl
3763	"4-(N,N-dimethylamino) phenylmethyl"	p-aminophenyl- methyl	n-butyl	propyl
3764	benzyl	"4-(N,N-dimethylamino)ph enylmethyl"	isobutyl	n-butyl
3765	1-pyrolylmethyl	benzyl	isobutyl	n-butyl

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3766	1-pyrazolylmethy l	1-pyrololymethyl	isobutyl	n-butyl
3767	1-imidazolylmeth yl	1-pyrazolylmethy l	isobutyl	n-butyl
3768	1-indolylmethyl	1-imidazolylmethy l	isobutyl	n-butyl
3769	1-triazolylmethy l	1-indolylmethyl	isobutyl	n-butyl
3770	1-tetrazolylmeth yl	1-triazolylmethy l	isobutyl	n-butyl
3771	2-pyridylmethyl	1-tetrazolylmethy l	isobutyl	n-butyl
3772	3-pyridylmethyl	2-pyridylmethyl	isobutyl	n-butyl
3773	4-pyridylmethyl	3-pyridylmethyl	isobutyl	n-butyl
3774	cyclohexylmeth yl	4-pyridylmethyl	isobutyl	n-butyl
3775	2-naphthylmethyl	cyclohexylmethyl	isobutyl	n-butyl
3776	3-naphthylmethyl	2-naphthylmethyl	isobutyl	n-butyl
3777	2-thiophenylmeth yl	3-naphthylmethyl	isobutyl	n-butyl
3778	4-(1-methyl)piperid inyl-methyl	2-thiophenylmethy l	isobutyl	n-butyl
3779	"(3,4-methylenedioxy phenyl)methyl"	4-(1-methyl)piperidin yl-methyl	isobutyl	n-butyl
3780	2-thienylmethyl	"(3,4-methylenedioxyph enyl)methyl"	isobutyl	n-butyl
3781	4-biphenylmethyl	2-thienylmethyl	isobutyl	n-butyl
3782	pyrimidinylmet hyl	4-biphenylmethyl	isobutyl	n-butyl
3783	2-benzothiazolyl methyl	pyrimidinylmethy l	isobutyl	n-butyl
3784	2-benzothiopheny lmethyl	2-benzothiazolylme thyl	isobutyl	n-butyl
3785	2-thiomethylethy l	2-benzothiophenylm ethyl	isobutyl	n-butyl
3786	2-thiomethylmeth yl	2-thiomethylethyl	isobutyl	n-butyl
3787	2-methylpropyl	2-thiomethylmethy l	isobutyl	n-butyl
3788	2-methylbutyl	2-methylpropyl	isobutyl	n-butyl

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3789	3-methylbutyl	2-methylbutyl	isobutyl	n-butyl
3790	cyclopropylmet hyl	3-methylbutyl	isobutyl	n-butyl
3791	cyclobutylmeth yl	cyclopropylmethy l	isobutyl	n-butyl
3792	cyclopentylmet hyl	cyclobutylmethyl	isobutyl	n-butyl
3793	p- hydroxyphenyl- methyl	cyclopentylmethy l	isobutyl	n-butyl
3794	p- nitrophenylmet hyl	p-hydroxyphenyl- methyl	isobutyl	n-butyl
3795	p-aminophenyl- methyl	p- nitrophenylmethy l	isobutyl	n-butyl
3796	"4-(N,N- dimethylamino) phenylmethyl" benzyl	p-aminophenyl- methyl	isobutyl	n-butyl
3797		"4-(N,N- dimethylamino)ph enylmethyl" benzyl	2-butyl	isobutyl
3798	1- pyrolylmethyl		2-butyl	isobutyl
3799	1- pyrazolylmethy l	1-pyrolylmethyl	2-butyl	isobutyl
3800	1- imidazolylmeth yl	1- pyrazolylmethyl	2-butyl	isobutyl
3801	1- indolylmethyl	1- imidazolylmethyl	2-butyl	isobutyl
3802	1- triazolylmethy l	1-indolylmethyl	2-butyl	isobutyl
3803	1- tetrazolylmeth yl	1- triazolylmethyl	2-butyl	isobutyl
3804	2- pyridylmethyl	1- tetrazolylmethyl	2-butyl	isobutyl
3805	3- pyridylmethyl	2-pyridylmethyl	2-butyl	isobutyl
3806	4- pyridylmethyl	3-pyridylmethyl	2-butyl	isobutyl
3807	cyclohexylmeth yl	4-pyridylmethyl	2-butyl	isobutyl
3808	2- naphthylmethyl	cyclohexylmethyl	2-butyl	isobutyl
3809	3- naphthylmethyl	2-naphthylmethyl	2-butyl	isobutyl
3810	2- thiophenylmeth yl	3-naphthylmethyl	2-butyl	isobutyl
3811	4-(1- methyl)piperid inyl-methyl	2- thiophenylmethyl	2-butyl	isobutyl

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3812	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-butyl	isobutyl
3813	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-butyl	isobutyl
3814	4-biphenylmethyl	2-thienylmethyl	2-butyl	isobutyl
3815	pyrimidinylmethyl	4-biphenylmethyl	2-butyl	isobutyl
3816	2-benzothiazolylmethyl	pyrimidinylmethyl	2-butyl	isobutyl
3817	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-butyl	isobutyl
3818	2-thiomethylethyl	2-benzothiophenylmethyl	2-butyl	isobutyl
3819	2-thiomethylmethyl	2-thiomethylethyl	2-butyl	isobutyl
3820	2-methylpropyl	2-thiomethylmethyl	2-butyl	isobutyl
3821	2-methylbutyl	2-methylpropyl	2-butyl	isobutyl
3822	3-methylbutyl	2-methylbutyl	2-butyl	isobutyl
3823	cyclopropylmethyl	3-methylbutyl	2-butyl	isobutyl
3824	cyclobutylmethyl	cyclopropylmethyl	2-butyl	isobutyl
3825	cyclopentylmethyl	cyclobutylmethyl	2-butyl	isobutyl
3826	p-hydroxyphenylmethyl	cyclopentylmethyl	2-butyl	isobutyl
3827	p-nitrophenylmethyl	p-hydroxyphenylmethyl	2-butyl	isobutyl
3828	p-aminophenylmethyl	p-nitrophenylmethyl	2-butyl	isobutyl
3829	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	2-butyl	isobutyl
3830	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	"3,3-dimethylallyl"	2-butyl
3831	1-pyrollylmethyl	benzyl	"3,3-dimethylallyl"	2-butyl
3832	1-pyrazolylmethyl	1-pyrollylmethyl	"3,3-dimethylallyl"	2-butyl
3833	1-imidazolylmethyl	1-pyrazolylmethyl	"3,3-dimethylallyl"	2-butyl
3834	1-indolylmethyl	1-imidazolylmethyl	"3,3-dimethylallyl"	2-butyl

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3835	1-triazolylmethyl	1-indolylmethyl	"3,3-dimethallyl"	2-butyl
3836	1-tetrazolylmethyl	1-triazolylmethyl	"3,3-dimethallyl"	2-butyl
3837	2-pyridylmethyl	1-tetrazolylmethyl	"3,3-dimethallyl"	2-butyl
3838	3-pyridylmethyl	2-pyridylmethyl	"3,3-dimethallyl"	2-butyl
3839	4-pyridylmethyl	3-pyridylmethyl	"3,3-dimethallyl"	2-butyl
3840	cyclohexylmethyl	4-pyridylmethyl	"3,3-dimethallyl"	2-butyl
3841	2-naphthylmethyl	cyclohexylmethyl	"3,3-dimethallyl"	2-butyl
3842	3-naphthylmethyl	2-naphthylmethyl	"3,3-dimethallyl"	2-butyl
3843	2-thiophenylmethyl	3-naphthylmethyl	"3,3-dimethallyl"	2-butyl
3844	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	"3,3-dimethallyl"	2-butyl
3845	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	"3,3-dimethallyl"	2-butyl
3846	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	"3,3-dimethallyl"	2-butyl
3847	4-biphenylmethyl	2-thienylmethyl	"3,3-dimethallyl"	2-butyl
3848	pyrimidinylmethyl	4-biphenylmethyl	"3,3-dimethallyl"	2-butyl
3849	2-benzothiazolylmethyl	pyrimidinylmethyl	"3,3-dimethallyl"	2-butyl
3850	2-benzothiophenylmethyl	2-benzothiazolylmethyl	"3,3-dimethallyl"	2-butyl
3851	2-thiomethylethyl	2-benzothiophenylethyl	"3,3-dimethallyl"	2-butyl
3852	2-thiomethylmethyl	2-thiomethylethyl	"3,3-dimethallyl"	2-butyl
3853	2-methylpropyl	2-thiomethylmethyl	"3,3-dimethallyl"	2-butyl
3854	2-methylbutyl	2-methylpropyl	"3,3-dimethallyl"	2-butyl
3855	3-methylbutyl	2-methylbutyl	"3,3-dimethallyl"	2-butyl
3856	cyclopropylmethyl	3-methylbutyl	"3,3-dimethallyl"	2-butyl
3857	cyclobutylmethyl	cyclopropylmethyl	"3,3-dimethallyl"	2-butyl

Table 6

3858	cyclopentylmeth hyl	cyclobutylmethyl	"3,3- dimethallyl"	2-butyl
3859	p- hydroxyphenyl- methyl	cyclopentylmethy l	"3,3- dimethallyl"	2-butyl
3860	p- nitrophenylmet hyl	p-hydroxyphenyl- methyl	"3,3- dimethallyl"	2-butyl
3861	p-aminophenyl- methyl	p- nitrophenylmethy l	"3,3- dimethallyl"	2-butyl
3862	"4-(N,N- dimethylamino) phenylmethyl"	p-aminophenyl- methyl	"3,3- dimethallyl"	2-butyl
3863	benzyl	"4-(N,N- dimethylamino)ph enylmethyl"	3-methallyl	"3,3- dimethallyl"
3864	1- pyrolylmethyl	benzyl	3-methallyl	"3,3- dimethallyl"
3865	1- pyrazolylmethy l	1-pyrolylmethyl	3-methallyl	"3,3- dimethallyl"
3866	1- imidazolylmeth yl	1- pyrazolylmethyl	3-methallyl	"3,3- dimethallyl"
3867	1- indolylmethyl	1- imidazolylmethyl	3-methallyl	"3,3- dimethallyl"
3868	1- triazolylmethy l	1-indolylmethyl	3-methallyl	"3,3- dimethallyl"
3869	1- tetrazolylmeth yl	1- triazolylmethyl	3-methallyl	"3,3- dimethallyl"
3870	2- pyridylmethyl	1- tetrazolylmethyl	3-methallyl	"3,3- dimethallyl"
3871	3- pyridylmethyl	2-pyridylmethyl	3-methallyl	"3,3- dimethallyl"
3872	4- pyridylmethyl	3-pyridylmethyl	3-methallyl	"3,3- dimethallyl"
3873	cyclohexylmeth yl	4-pyridylmethyl	3-methallyl	"3,3- dimethallyl"
3874	2- naphthylmethyl	cyclohexylmethyl	3-methallyl	"3,3- dimethallyl"
3875	3- naphthylmethyl	2-naphthylmethyl	3-methallyl	"3,3- dimethallyl"
3876	2- thiophenylmeth yl	3-naphthylmethyl	3-methallyl	"3,3- dimethallyl"
3877	4-(1- methyl)piperid inyl-methyl	2- thiophenylmethyl	3-methallyl	"3,3- dimethallyl"
3878	"(3,4- methylenedioxy phenyl)methyl"	4-(1- methyl)piperidin yl-methyl	3-methallyl	"3,3- dimethallyl"
3879	2- thienylmethyl	"(3,4- methylenedioxyph enyl)methyl"	3-methallyl	"3,3- dimethallyl"

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3880	4-biphenylmethyl	2-thienylmethyl	3-methallyl	"3,3-dimethallyl"
3881	pyrimidinyl-methyl	4-biphenylmethyl	3-methallyl	"3,3-dimethallyl"
3882	2-benzothiazolylmethyl	pyrimidinylmethy l	3-methallyl	"3,3-dimethallyl"
3883	2-benzothio-phenylmethyl	2-benzothiazolylme thyl	3-methallyl	"3,3-dimethallyl"
3884	2-thiomethyl-ethyl	2-benzothiophenylm ethyl	3-methallyl	"3,3-dimethallyl"
3885	2-thiomethyl-methyl	2-thiomethylethyl	3-methallyl	"3,3-dimethallyl"
3886	2-methylpropyl	2-thiomethylmethyl	3-methallyl	"3,3-dimethallyl"
3887	2-methylbutyl	2-methylpropyl	3-methallyl	"3,3-dimethallyl"
3888	3-methylbutyl	2-methylbutyl	3-methallyl	"3,3-dimethallyl"
3889	cyclopropylmet hyl	3-methylbutyl	3-methallyl	"3,3-dimethallyl"
3890	cyclobutylmeth yl	cyclopropylmethy l	3-methallyl	"3,3-dimethallyl"
3891	cyclopentylmet hyl	cyclobutylmethyl	3-methallyl	"3,3-dimethallyl"
3892	p-hydroxyphenyl-methyl	cyclopentylmethy l	3-methallyl	"3,3-dimethallyl"
3893	p-nitrophenylmet hyl	p-hydroxyphenyl-methyl	3-methallyl	"3,3-dimethallyl"
3894	p-aminophenyl-methyl	p-nitrophenylmethy l	3-methallyl	"3,3-dimethallyl"
3895	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	3-methallyl	"3,3-dimethallyl"
3896	benzyl	"4-(N,N-dimethylamino)ph enylmethyl"	2-methallyl	3-methallyl
3897	1-pyrollylmethyl	benzyl	2-methallyl	3-methallyl
3898	1-pyrazolyl-methyl	1-pyrollylmethyl	2-methallyl	3-methallyl
3899	1-imidazolyl-methyl	1-pyrazolylmethyl	2-methallyl	3-methallyl
3900	1-indolylmethyl	1-imidazolylmethyl	2-methallyl	3-methallyl
3901	1-triazolyl-methyl	1-indolylmethyl	2-methallyl	3-methallyl
3902	1-tetrazolyl-methyl	1-triazolylmethyl	2-methallyl	3-methallyl
3903	2-pyridylmethyl	1-tetrazolylmethyl	2-methallyl	3-methallyl

Table 6

3904	3-pyridylmethyl	2-pyridylmethyl	2-methallyl	3-methallyl
3905	4-pyridylmethyl	3-pyridylmethyl	2-methallyl	3-methallyl
3906	cyclohexylmethyl	4-pyridylmethyl	2-methallyl	3-methallyl
3907	2-naphthylmethyl	cyclohexylmethyl	2-methallyl	3-methallyl
3908	3-naphthylmethyl	2-naphthylmethyl	2-methallyl	3-methallyl
3909	2-thiophenylmethyl	3-naphthylmethyl	2-methallyl	3-methallyl
3910	4-(1-methyl)-piperidinylmethyl	2-thiophenylmethyl	2-methallyl	3-methallyl
3911	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinylmethyl	2-methallyl	3-methallyl
3912	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-methallyl	3-methallyl
3913	4-biphenylmethyl	2-thienylmethyl	2-methallyl	3-methallyl
3914	pyrimidinylmethyl	4-biphenylmethyl	2-methallyl	3-methallyl
3915	2-benzothiazolylmethyl	pyrimidinylmethyl	2-methallyl	3-methallyl
3916	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-methallyl	3-methallyl
3917	2-thiomethylethyl	2-benzothiophenylmethyl	2-methallyl	3-methallyl
3918	2-thiomethylmethyl	2-thiomethylethyl	2-methallyl	3-methallyl
3919	2-methylpropyl	2-thiomethylmethyl	2-methallyl	3-methallyl
3920	2-methylbutyl	2-methylpropyl	2-methallyl	3-methallyl
3921	3-methylbutyl	2-methylbutyl	2-methallyl	3-methallyl
3922	cyclopropylmethyl	3-methylbutyl	2-methallyl	3-methallyl
3923	cyclobutylmethyl	cyclopropylmethyl	2-methallyl	3-methallyl
3924	cyclopentylmethyl	cyclobutylmethyl	2-methallyl	3-methallyl
3925	p-hydroxyphenylmethyl	cyclopentylmethyl	2-methallyl	3-methallyl
3926	p-nitrophenylmethyl	p-hydroxyphenylmethyl	2-methallyl	3-methallyl
3927	p-aminophenylmethyl	p-nitrophenylmethyl	2-methallyl	3-methallyl
3928	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	2-methallyl	3-methallyl

Table 6

3929	benzyl	"4-(N,N-dimethylamino)-phenylmethyl"	2-propyl	2-methallyl
3930	1-pyrolylmethyl	benzyl	2-propyl	2-methallyl
3931	1-pyrazolyl-methyl	1-pyrolylmethyl	2-propyl	2-methallyl
3932	1-imidazolyl-methyl	1-pyrazolylmethyl	2-propyl	2-methallyl
3933	1-indolylmethyl	1-imidazolylmethyl	2-propyl	2-methallyl
3934	1-triazolylmethy	1-indolylmethyl	2-propyl	2-methallyl
3935	1-tetrazolylmeth	1-triazolylmethyl	2-propyl	2-methallyl
3936	2-pyridylmethyl	1-tetrazolylmethy	2-propyl	2-methallyl
3937	3-pyridylmethyl	2-pyridylmethyl	2-propyl	2-methallyl
3938	4-pyridylmethyl	3-pyridylmethyl	2-propyl	2-methallyl
3939	pyridylmethyl	4-pyridylmethyl	2-propyl	2-methallyl
3940	2-cyclohexyl-methyl	cyclohexylmethyl	2-propyl	2-methallyl
3941	3-naphthylmethyl	2-naphthylmethyl	2-propyl	2-methallyl
3942	2-naphthylmethyl	3-naphthylmethyl	2-propyl	2-methallyl
3943	4-(1-methyl)-piperidinyl-methyl	2-thiophenylmethyl	2-propyl	2-methallyl
3944	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)-piperidinyl-methyl	2-propyl	2-methallyl
3945	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-propyl	2-methallyl
3946	4-biphenylmethyl	2-thienylmethyl	2-propyl	2-methallyl
3947	pyrimidinyl-methyl	4-biphenylmethyl	2-propyl	2-methallyl
3948	2-benzothiazolyl methyl	pyrimidinyl-methyl	2-propyl	2-methallyl
3949	2-benzothio-phenylmethyl	2-benzothiazolyl-methyl	2-propyl	2-methallyl
3950	2-thio-methylethyl	2-benzothiophenyl-methyl	2-propyl	2-methallyl
3951	2-thio-methylmethyl	2-thiomethylethyl	2-propyl	2-methallyl
3952	2-methylpropyl	2-thiomethylmethyl	2-propyl	2-methallyl

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3953	2-methylbutyl	2-methylpropyl	2-propyl	2-methallyl
3954	3-methylbutyl	2-methylbutyl	2-propyl	2-methallyl
3955	cyclopropylmet hyl	3-methylbutyl	2-propyl	2-methallyl
3956	cyclobutylmeth yl	cyclopropylmethy l	2-propyl	2-methallyl
3957	cyclopentylmet hyl	cyclobutylmethy l	2-propyl	2-methallyl
3958	p- hydroxyphenyl- methyl	cyclopentylmethy l	2-propyl	2-methallyl
3959	p- nitrophenylmet hyl	p-hydroxyphenyl- methyl	2-propyl	2-methallyl
3960	p-aminophenyl- methyl	p-nitrophenyl- methyl	2-propyl	2-methallyl
3961	"4-(N,N- dimethylamino) phenylmethyl" benzyl	p-aminophenyl- methyl	2-propyl	2-methallyl
3962		"4-(N,N- dimethylamino)- phenylmethyl" benzyl	cyclopropyl	2-propyl
3963	1- pyrolylmethyl		cyclopropyl	2-propyl
3964	1-pyrazolyl- methyl	1-pyrolylmethyl	cyclopropyl	2-propyl
3965	1-imidazolyl- methyl	1- pyrazolylmethyl	cyclopropyl	2-propyl
3966	1- indolylmethyl	1- imidazolylmethyl	cyclopropyl	2-propyl
3967	1-triazolyl- methyl	1-indolylmethyl	cyclopropyl	2-propyl
3968	1-tetrazolyl- methyl	1- triazolylmethyl	cyclopropyl	2-propyl
3969	2- pyridylmethyl	1- tetrazolylmethyl	cyclopropyl	2-propyl
3970	3- pyridylmethyl	2-pyridylmethyl	cyclopropyl	2-propyl
3971	4- pyridylmethyl	3-pyridylmethyl	cyclopropyl	2-propyl
3972	cyclohexyl- methyl	4-pyridylmethyl	cyclopropyl	2-propyl
3973	2- naphthylmethyl	cyclohexylmethyl	cyclopropyl	2-propyl
3974	3- naphthylmethyl	2-naphthylmethyl	cyclopropyl	2-propyl
3975	2- thiophenylmeth yl	3-naphthylmethyl	cyclopropyl	2-propyl
3976	4-(1- methyl)piperid inyl-methyl	2- thiophenylmethyl	cyclopropyl	2-propyl
3977	"(3,4- methylenedioxy phenyl)methyl"	4-(1-methyl)- piperidinyl- methyl	cyclopropyl	2-propyl

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3978	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclopropyl	2-propyl
3979	4-biphenylmethyl	2-thienylmethyl	cyclopropyl	2-propyl
3980	pyrimidinyl-methyl	4-biphenylmethyl	cyclopropyl	2-propyl
3981	2-benzothiazolylmethyl	pyrimidinyl-methyl	cyclopropyl	2-propyl
3982	2-benzothio-phenylmethyl	2-benzothiazolyl-methyl	cyclopropyl	2-propyl
3983	2-thio-methylethyl	2-benzothiophenylmethyl	cyclopropyl	2-propyl
3984	2-thio-methylmethyl	2-thiomethylethyl	cyclopropyl	2-propyl
3985	2-methylpropyl	2-thiomethylmethyl	cyclopropyl	2-propyl
3986	2-methylbutyl	2-methylpropyl	cyclopropyl	2-propyl
3987	3-methylbutyl	2-methylbutyl	cyclopropyl	2-propyl
3988	cyclopropylmethyl	3-methylbutyl	cyclopropyl	2-propyl
3989	cyclobutylmethyl	cyclopropyl-methyl	cyclopropyl	2-propyl
3990	cyclopentylmethyl	cyclobutylmethyl	cyclopropyl	2-propyl
3991	p-hydroxyphenyl-methyl	cyclopentylmethyl	cyclopropyl	2-propyl
3992	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	cyclopropyl	2-propyl
3993	p-aminophenyl-methyl	p-nitrophenylmethyl	cyclopropyl	2-propyl
3994	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	cyclopropyl	2-propyl
3995	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	cyclopropyl-methyl	cyclopropyl
3996	1-pyrollylmethyl	benzyl	cyclopropyl-methyl	cyclopropyl
3997	1-pyrazolyl-methyl	1-pyrollylmethyl	cyclopropyl-methyl	cyclopropyl
3998	1-imidazolyl-methyl	1-pyrazolylmethyl	cyclopropyl-methyl	cyclopropyl
3999	1-indolylmethyl	1-imidazolylmethyl	cyclopropyl-methyl	cyclopropyl
4000	1-triazolyl-methyl	1-indolylmethyl	cyclopropyl-methyl	cyclopropyl
4001	1-tetrazolyl-methyl	1-triazolylmethyl	cyclopropyl-methyl	cyclopropyl
4002	2-pyridylmethyl	1-tetrazolylmethyl	cyclopropyl-methyl	cyclopropyl

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4003	3-pyridylmethyl	2-pyridylmethyl	cyclopropyl-methyl-	cyclopropyl
4004	4-pyridylmethyl	3-pyridylmethyl	cyclopropylmet	cyclopropyl
4005	cyclohexyl-methyl	4-pyridylmethyl	hyl	cyclopropyl
4006	2-naphthylmethyl	cyclohexylmethyl	cyclopropyl-methyl	cyclopropyl
4007	3-naphthylmethyl	2-naphthylmethyl	cyclopropylmet	cyclopropyl
4008	2-thiophenyl-methyl	3-naphthylmethyl	hyl	cyclopropyl
4009	4-(1-methyl)-piperidinyl-methyl	2-thiophenyl-methyl	cyclopropyl-methyl	cyclopropyl
4010	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)-piperidinyl-methyl	cyclopropyl-methyl	cyclopropyl
4011	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclopropyl-methyl	cyclopropyl
4012	4-biphenylmethyl	2-thienylmethyl	cyclopropyl-methyl	cyclopropyl
4013	pyrimidinyl-methyl	4-biphenylmethyl	cyclopropyl-methyl	cyclopropyl
4014	2-benzothiazolyl-methyl	pyrimidinylmethy	cyclopropyl-methyl	cyclopropyl
4015	2-benzothio-phenylmethyl	l		
4016	2-thiomethyl-ethyl	2-benzo-thiazolyl-methyl	cyclopropyl-methyl	cyclopropyl
4017	2-thiomethylmeth	2-benzo-thiophenylmethyl	cyclopropyl-methyl	cyclopropyl
4018	yl	2-thiomethylethyl	cyclopropyl-methyl	cyclopropyl
4019	2-methylpropyl	2-thiomethylmethyl	cyclopropyl-methyl	cyclopropyl
4020	2-methylbutyl	2-methylpropyl	cyclopropyl-methyl	cyclopropyl
4021	3-methylbutyl	2-methylbutyl	cyclopropyl-methyl	cyclopropyl
4022	cyclopropyl-methyl	3-methylbutyl	cyclopropyl-methyl	cyclopropyl
4023	cyclobutyl-methyl	cyclopropyl-methyl	cyclopropyl-methyl	cyclopropyl
4024	cyclopentyl-methyl	cyclobutyl-methyl	cyclopropyl-methyl	cyclopropyl
4025	p-hydroxyphenyl-methyl	cyclopentyl-methyl	cyclopropyl-methyl	cyclopropyl
4026	p-nitrophenyl-methyl	p-hydroxyphenyl-methyl	cyclopropylmet	cyclopropyl
4027	p-aminophenyl-methyl	p-nitrophenyl-methyl	hyl	cyclopropyl
	"4-(N,N-di-methyl-amino)-phenyl-methyl"	p-aminophenyl-methyl	cyclopropyl-methyl	cyclopropyl
			cyclopropyl-methyl	cyclopropyl

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4028	benzyl	"4-(N,N-dimethylamino)-phenylmethyl"	n-pentyl	cyclopropylmeth yl
4029	1-pyrolylmethyl	benzyl	n-pentyl	cyclopropylmeth yl
4030	1-pyrazolylmethy l	1-pyrolylmethyl	n-pentyl	cyclopropylmeth yl
4031	1-imidazolylmeth yl	1-pyrazolylmethy l	n-pentyl	cyclopropylmeth yl
4032	1-indolylmethyl	1-imidazolylmethy l	n-pentyl	cyclopropylmeth yl
4033	1-triazolylmethy l	1-indolylmethyl	n-pentyl	cyclopropylmeth yl
4034	1-tetrazolylmeth yl	1-triazolylmethyl	n-pentyl	cyclopropylmeth yl
4035	2-pyridylmethyl	1-tetrazolylmethy l	n-pentyl	cyclopropylmeth yl
4036	3-pyridylmethyl	2-pyridylmethyl	n-pentyl	cyclopropylmeth yl
4037	4-pyridylmethyl	3-pyridylmethyl	n-pentyl	cyclopropylmeth yl
4038	cyclohexylmeth yl	4-pyridylmethyl	n-pentyl	cyclopropylmeth yl
4039	2-naphthylmethyl	cyclohexylmethyl	n-pentyl	cyclopropylmeth yl
4040	3-naphthylmethyl	2-naphthylmethyl	n-pentyl	cyclopropylmeth yl
4041	2-thiophenylmeth yl	3-naphthylmethyl	n-pentyl	cyclopropylmeth yl
4042	4-(1-methyl)piperid inyl-methyl	2-thiophenylmethy l	n-pentyl	cyclopropylmeth yl
4043	"(3,4-methylenedioxy phenyl)methyl"	4-(1-methyl)piperidin yl-methyl	n-pentyl	cyclopropylmeth yl
4044	2-thienylmethyl	"(3,4-methylenedioxyph enyl)methyl"	n-pentyl	cyclopropylmeth yl
4045	4-biphenylmethyl	2-thienylmethyl	n-pentyl	cyclopropylmeth yl
4046	pyrimidinylmet hyl	4-biphenylmethyl	n-pentyl	cyclopropylmeth yl
4047	2-benzothiazolyl methyl	pyrimidinylmethy l	n-pentyl	cyclopropylmeth yl
4048	2-benzothiopheny lmethyl	2-benzothiazolylme thyl	n-pentyl	cyclopropylmeth yl
4049	2-thiomethylethy l	2-benzothiophenylm ethyl	n-pentyl	cyclopropylmeth yl

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4050	2-thiomethylmeth yl	2-thiomethylethyl	n-pentyl	cyclopropylmeth yl
4051	2-methylpropyl	2-thiomethylmeth yl	n-pentyl	cyclopropylmeth yl
4052	2-methylbutyl	2-methylpropyl	n-pentyl	cyclopropylmeth yl
4053	3-methylbutyl	2-methylbutyl	n-pentyl	cyclopropylmeth yl
4054	cyclopropylmet hyl	3-methylbutyl	n-pentyl	cyclopropylmeth yl
4055	cyclobutylmeth yl	cyclopropylmeth yl	n-pentyl	cyclopropylmeth yl
4056	cyclopentylmet hyl	cyclobutylmeth yl	n-pentyl	cyclopropylmeth yl
4057	p-hydroxyphenyl- methyl	cyclopentylmeth yl	n-pentyl	cyclopropylmeth yl
4058	p-nitrophenylmet hyl	p-hydroxyphenyl- methyl	n-pentyl	cyclopropylmeth yl
4059	p-aminophenyl- methyl	p-nitrophenylmeth yl	n-pentyl	cyclopropylmeth yl
4060	"4-(N,N- dimethylamino) phenylmethyl" benzyl	p-aminophenyl- methyl	n-pentyl	cyclopropylmeth yl
4061		"4-(N,N- dimethylamino)ph enylmethyl" benzyl	2-pentyl	n-pentyl
4062	1-pyrollylmethyl		2-pentyl	n-pentyl
4063	1-pyrazolylmeth yl	1-pyrollylmethyl	2-pentyl	n-pentyl
4064	1-imidazolylmeth yl	1-pyrazolylmeth yl	2-pentyl	n-pentyl
4065	1-indolylmethyl	1-imidazolylmeth yl	2-pentyl	n-pentyl
4066	1-triazolylmeth yl	1-indolylmethyl	2-pentyl	n-pentyl
4067	1-tetrazolylmeth yl	1-triazolylmeth yl	2-pentyl	n-pentyl
4068	2-pyridylmethyl	1-tetrazolylmeth yl	2-pentyl	n-pentyl
4069	3-pyridylmethyl	2-pyridylmethyl	2-pentyl	n-pentyl
4070	4-pyridylmethyl	3-pyridylmethyl	2-pentyl	n-pentyl
4071	cyclohexylmeth yl	4-pyridylmethyl	2-pentyl	n-pentyl
4072	2-naphthylmeth yl	cyclohexylmeth yl	2-pentyl	n-pentyl

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4073	3-naphthylmethyl	2-naphthylmethyl	2-pentyl	n-pentyl
4074	2-thiophenylmethyl	3-naphthylmethyl	2-pentyl	n-pentyl
4075	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-pentyl	n-pentyl
4076	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-pentyl	n-pentyl
4077	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-pentyl	n-pentyl
4078	4-biphenylmethyl	2-thienylmethyl	2-pentyl	n-pentyl
4079	pyrimidinylmethyl	4-biphenylmethyl	2-pentyl	n-pentyl
4080	2-benzothiazolylmethyl	pyrimidinylmethyl	2-pentyl	n-pentyl
4081	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-pentyl	n-pentyl
4082	2-thiomethylethyl	2-benzothiophenylmethyl	2-pentyl	n-pentyl
4083	2-thiomethylmethyl	2-thiomethylethyl	2-pentyl	n-pentyl
4084	2-methylpropyl	2-thiomethylmethyl	2-pentyl	n-pentyl
4085	2-methylbutyl	2-methylpropyl	2-pentyl	n-pentyl
4086	3-methylbutyl	2-methylbutyl	2-pentyl	n-pentyl
4087	cyclopropylmethyl	3-methylbutyl	2-pentyl	n-pentyl
4088	cyclobutylmethyl	cyclopropylmethyl	2-pentyl	n-pentyl
4089	cyclopentylmethyl	cyclobutylmethyl	2-pentyl	n-pentyl
4090	p-hydroxyphenylmethyl	cyclopentylmethyl	2-pentyl	n-pentyl
4091	p-nitrophenylmethyl	p-hydroxyphenylmethyl	2-pentyl	n-pentyl
4092	p-aminophenylmethyl	p-nitrophenylmethyl	2-pentyl	n-pentyl
4093	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	2-pentyl	n-pentyl
4094	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	3-pentyl	2-pentyl
4095	1-pyrollylmethyl	benzyl	3-pentyl	2-pentyl

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4096	1-pyrazolylmethy l	1-pyrazolylmethy l	3-pentyl	2-pentyl
4097	1-imidazolylmeth yl	1-pyrazolylmethy l	3-pentyl	2-pentyl
4098	1-indolylmethy l	1-imidazolylmethy l	3-pentyl	2-pentyl
4099	1-triazolylmethy l	1-indolylmethy l	3-pentyl	2-pentyl
4100	1-tetrazolylmeth yl	1-triazolylmethy l	3-pentyl	2-pentyl
4101	2-pyridylmethy l	1-tetrazolylmethy l	3-pentyl	2-pentyl
4102	3-pyridylmethy l	2-pyridylmethy l	3-pentyl	2-pentyl
4103	4-pyridylmethy l	3-pyridylmethy l	3-pentyl	2-pentyl
4104	cyclohexylmeth yl	4-pyridylmethy l	3-pentyl	2-pentyl
4105	2-naphthylmethy l	cyclohexylmethy l	3-pentyl	2-pentyl
4106	3-naphthylmethy l	2-naphthylmethy l	3-pentyl	2-pentyl
4107	2-thiophenylmeth yl	3-naphthylmethy l	3-pentyl	2-pentyl
4108	4-(1-methyl)piperid inyl-methyl	2-thiophenylmethy l	3-pentyl	2-pentyl
4109	"(3,4-methylenedioxy phenyl)methyl"	4-(1-methyl)piperidin yl-methyl	3-pentyl	2-pentyl
4110	2-thienylmethy l	"(3,4-methylenedioxyph enyl)methyl"	3-pentyl	2-pentyl
4111	4-biphenylmethy l	2-thienylmethy l	3-pentyl	2-pentyl
4112	pyrimidinylmet hyl	4-biphenylmethy l	3-pentyl	2-pentyl
4113	2-benzothiazolyl methyl	pyrimidinylmethy l	3-pentyl	2-pentyl
4114	2-benzothiopheny lmethyl	2-benzothiazolylme thyl	3-pentyl	2-pentyl
4115	2-thiomethylethy l	2-benzothiophenylm ethyl	3-pentyl	2-pentyl
4116	2-thiomethylmeth yl	2-thiomethylethy l	3-pentyl	2-pentyl
4117	2-methylpropyl	2-thiomethylmethy l	3-pentyl	2-pentyl
4118	2-methylbutyl	2-methylpropyl	3-pentyl	2-pentyl

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4119	3-methylbutyl	2-methylbutyl	3-pentyl	2-pentyl
4120	cyclopropylmeth hyl	3-methylbutyl	3-pentyl	2-pentyl
4121	cyclobutylmeth yl	cyclopropylmethy l	3-pentyl	2-pentyl
4122	cyclopentylmet hyl	cyclobutylmethy l	3-pentyl	2-pentyl
4123	p- hydroxyphenyl- methyl	cyclopentylmethy l	3-pentyl	2-pentyl
4124	p- nitrophenylmet hyl	p-hydroxyphenyl- methyl	3-pentyl	2-pentyl
4125	p-aminophenyl- methyl	p- nitrophenylmethy l	3-pentyl	2-pentyl
4126	"4-(N,N- dimethylamino) phenylmethyl" benzyl	p-aminophenyl- methyl	3-pentyl	2-pentyl
4127		"4-(N,N- dimethylamino)ph enylmethyl" benzyl	3-methylbutyl	3-pentyl
4128	1- pyrolylmethyl		3-methylbutyl	3-pentyl
4129	1- pyrazolylmethy l	1-pyrolylmethyl	3-methylbutyl	3-pentyl
4130	1- imidazolylmeth yl	1- pyrazolylmethy l	3-methylbutyl	3-pentyl
4131	1- indolylmethyl	1- imidazolylmethy l	3-methylbutyl	3-pentyl
4132	1- triazolylmethy l	1-indolylmethyl	3-methylbutyl	3-pentyl
4133	1- tetrazolylmeth yl	1- triazolylmethy l	3-methylbutyl	3-pentyl
4134	2- pyridylmethyl	1- tetrazolylmethy l	3-methylbutyl	3-pentyl
4135	3- pyridylmethyl	2-pyridylmethyl	3-methylbutyl	3-pentyl
4136	4- pyridylmethyl	3-pyridylmethyl	3-methylbutyl	3-pentyl
4137	cyclohexylmeth yl	4-pyridylmethyl	3-methylbutyl	3-pentyl
4138	2- naphthylmethy l	cyclohexylmethy l	3-methylbutyl	3-pentyl
4139	3- naphthylmethy l	2-naphthylmethy l	3-methylbutyl	3-pentyl
4140	2- thiophenylmeth yl	3-naphthylmethy l	3-methylbutyl	3-pentyl
4141	4-(1- methyl)piperid inyl-methyl	2- thiophenylmethy l	3-methylbutyl	3-pentyl

Table 6

4142	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	3-methylbutyl	3-pentyl
4143	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-methylbutyl	3-pentyl
4144	4-biphenylmethyl	2-thienylmethyl	3-methylbutyl	3-pentyl
4145	pyrimidinylmethyl	4-biphenylmethyl	3-methylbutyl	3-pentyl
4146	2-benzothiazolylmethyl	pyrimidinylmethyl	3-methylbutyl	3-pentyl
4147	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-methylbutyl	3-pentyl
4148	2-thiomethylethyl	2-benzothiophenylmethyl	3-methylbutyl	3-pentyl
4149	2-thiomethylmethyl	2-thiomethylethyl	3-methylbutyl	3-pentyl
4150	2-methylpropyl	2-thiomethylmethyl	3-methylbutyl	3-pentyl
4151	2-methylbutyl	2-methylpropyl	3-methylbutyl	3-pentyl
4152	3-methylbutyl	2-methylbutyl	3-methylbutyl	3-pentyl
4153	cyclopropylmethyl	3-methylbutyl	3-methylbutyl	3-pentyl
4154	cyclobutylmethyl	cyclopropylmethyl	3-methylbutyl	3-pentyl
4155	cyclopentylmethyl	cyclobutylmethyl	3-methylbutyl	3-pentyl
4156	p-hydroxyphenylmethyl	cyclopentylmethyl	3-methylbutyl	3-pentyl
4157	p-nitrophenylmethyl	1-p-hydroxyphenylmethyl	3-methylbutyl	3-pentyl
4158	p-aminophenylmethyl	p-nitrophenylmethyl	3-methylbutyl	3-pentyl
4159	"4-(N,N-dimethylamino)phenylmethyl"	1-p-aminophenylmethyl	3-methylbutyl	3-pentyl
4160	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-methylbutyl	3-methylbutyl
4161	1-pyrollylmethyl	benzyl	2-methylbutyl	3-methylbutyl
4162	1-pyrazolylmethyl	1-pyrollylmethyl	2-methylbutyl	3-methylbutyl
4163	1-imidazolylmethyl	1-pyrazolylmethyl	2-methylbutyl	3-methylbutyl
4164	1-indolylmethyl	1-imidazolylmethyl	2-methylbutyl	3-methylbutyl

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4165	1- triazolylmethy l	1-indolylmethyl	2-methylbutyl	3-methylbutyl
4166	1- tetrazolylmeth yl	1- triazolylmethyl	2-methylbutyl	3-methylbutyl
4167	2- pyridylmethyl	1- tetrazolylmethyl	2-methylbutyl	3-methylbutyl
4168	3- pyridylmethyl	2-pyridylmethyl	2-methylbutyl	3-methylbutyl
4169	4- pyridylmethyl	3-pyridylmethyl	2-methylbutyl	3-methylbutyl
4170	cyclohexylmeth yl	4-pyridylmethyl	2-methylbutyl	3-methylbutyl
4171	2- naphthylmethyl	cyclohexylmethyl	2-methylbutyl	3-methylbutyl
4172	3- naphthylmethyl	2-naphthylmethyl	2-methylbutyl	3-methylbutyl
4173	2- thiophenylmeth yl	3-naphthylmethyl	2-methylbutyl	3-methylbutyl
4174	4-(1- methyl)piperid inyl-methyl	2- thiophenylmethyl	2-methylbutyl	3-methylbutyl
4175	"(3,4- methylenedioxy phenyl)methyl"	4-(1- methyl)piperidin yl-methyl	2-methylbutyl	3-methylbutyl
4176	2- thienylmethyl	"(3,4- methylenedioxyph enyl)methyl"	2-methylbutyl	3-methylbutyl
4177	4- biphenylmethyl	2-thienylmethyl	2-methylbutyl	3-methylbutyl
4178	pyrimidinylmet hyl	4-biphenylmethyl	2-methylbutyl	3-methylbutyl
4179	2- benzothiazolyl methyl	pyrimidinylmethy l	2-methylbutyl	3-methylbutyl
4180	2- benzothiopheny lmethyl	2- benzothiazolylme thyl	2-methylbutyl	3-methylbutyl
4181	2- thiomethylethy l	2- benzothiophenylm ethyl	2-methylbutyl	3-methylbutyl
4182	2- thiomethylmeth yl	2- thiomethylethyl	2-methylbutyl	3-methylbutyl
4183	2-methylpropyl	2- thiomethylmethyl	2-methylbutyl	3-methylbutyl
4184	2-methylbutyl	2-methylpropyl	2-methylbutyl	3-methylbutyl
4185	3-methylbutyl	2-methylbutyl	2-methylbutyl	3-methylbutyl
4186	cyclopropylmet hyl	3-methylbutyl	2-methylbutyl	3-methylbutyl
4187	cyclobutylmeth yl	cyclopropylmethy l	2-methylbutyl	3-methylbutyl
4188	cyclopentylmet hyl	cyclobutylmethyl	2-methylbutyl	3-methylbutyl

Table 6

4189	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-methylbutyl	3-methylbutyl
4190	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-methylbutyl	3-methylbutyl
4191	p-aminophenyl-methyl	p-nitrophenylmethyl	2-methylbutyl	3-methylbutyl
4192	"4-(N,N-dimethylamino)phenylmethyl" benzyl	p-aminophenyl-methyl	2-methylbutyl	3-methylbutyl
4193		"4-(N,N-dimethylamino)phenylmethyl" benzyl	propargyl	2-methylbutyl
4194	1-pyrolylmethyl		propargyl	2-methylbutyl
4195	1-pyrazolylmethyl	1-pyrolylmethyl	propargyl	2-methylbutyl
4196	1-imidazolylmethyl	1-pyrazolylmethyl	propargyl	2-methylbutyl
4197	1-indolylmethyl	1-imidazolylmethyl	propargyl	2-methylbutyl
4198	1-triazolylmethyl	1-indolylmethyl	propargyl	2-methylbutyl
4199	1-tetrazolylmethyl	1-triazolylmethyl	propargyl	2-methylbutyl
4200	2-pyridylmethyl	1-tetrazolylmethyl	propargyl	2-methylbutyl
4201	3-pyridylmethyl	2-pyridylmethyl	propargyl	2-methylbutyl
4202	4-pyridylmethyl	3-pyridylmethyl	propargyl	2-methylbutyl
4203	pyridylmethyl	4-pyridylmethyl	propargyl	2-methylbutyl
4204	cyclohexylmethyl	cyclohexylmethyl	propargyl	2-methylbutyl
4205	2-naphthylmethyl	2-naphthylmethyl	propargyl	2-methylbutyl
4206	3-naphthylmethyl	3-naphthylmethyl	propargyl	2-methylbutyl
4207	2-thiophenylmethyl	2-thiophenylmethyl	propargyl	2-methylbutyl
4208	4-(1-methyl)piperidinyl-methyl	4-(1-methyl)piperidinyl-methyl	propargyl	2-methylbutyl
4209	"(3,4-methylenedioxyphenyl)methyl"	"(3,4-methylenedioxyphenyl)methyl"	propargyl	2-methylbutyl
4210	2-thienylmethyl	2-thienylmethyl	propargyl	2-methylbutyl

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4211	pyrimidinylmet hyl	4-biphenylmethyl	propargyl	2-methylbutyl
4212	2- benzothiazolyl methyl	pyrimidinylmethy l	propargyl	2-methylbutyl
4213	2- benzothiopheny lmethyl	2- benzothiazolylme thyl	propargyl	2-methylbutyl
4214	2- thiomethylethy l	2- benzothiophenylm ethyl	propargyl	2-methylbutyl
4215	2- thiomethylmeth yl	2- thiomethylethyl	propargyl	2-methylbutyl
4216	2-methylpropyl	2- thiomethylmethyl	propargyl	2-methylbutyl
4217	2-methylbutyl	2-methylpropyl	propargyl	2-methylbutyl
4218	3-methylbutyl	2-methylbutyl	propargyl	2-methylbutyl
4219	cyclopropylmet hyl	3-methylbutyl	propargyl	2-methylbutyl
4220	cyclobutylmeth yl	cyclopropylmethy l	propargyl	2-methylbutyl
4221	cyclopentylmet hyl	cyclobutylmethyl	propargyl	2-methylbutyl
4222	p- hydroxyphenyl- methyl	cyclopentylmethy l	propargyl	2-methylbutyl
4223	p- nitrophenylmet hyl	p-hydroxyphenyl- methyl	propargyl	2-methylbutyl
4224	p-aminophenyl- methyl	p- nitrophenylmethy l	propargyl	2-methylbutyl
4225	"4-(N,N- dimethylamino) phenylmethyl" benzyl	p-aminophenyl- methyl	propargyl	2-methylbutyl
4226	1- pyrolylmethyl	"4-(N,N- dimethylamino)ph enylmethyl" benzyl	cyclobutyl	propargyl
4227	1- pyrazolylmethy l	1-pyrolylmethyl	cyclobutyl	propargyl
4228	1- imidazolylmeth yl	1- pyrazolylmethyl	cyclobutyl	propargyl
4229	1- indolylmethyl	1- imidazolylmethyl	cyclobutyl	propargyl
4230	1- triazolylmethy l	1-indolylmethyl	cyclobutyl	propargyl
4231	1- tetrazolylmeth yl	1- triazolylmethyl	cyclobutyl	propargyl
4232	2- pyridylmethyl	1- tetrazolylmethyl	cyclobutyl	propargyl

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4234	3-pyridylmethyl	2-pyridylmethyl	cyclobutyl	propargyl
4235	4-pyridylmethyl	3-pyridylmethyl	cyclobutyl	propargyl
4236	cyclohexylmeth	4-pyridylmethyl	cyclobutyl	propargyl
4237	yl	cyclohexylmethyl	cyclobutyl	propargyl
4238	2-naphthylmethyl	2-naphthylmethyl	cyclobutyl	propargyl
4239	3-naphthylmethyl	3-naphthylmethyl	cyclobutyl	propargyl
4240	2-thiophenylmeth			
4241	yl			
4242	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	cyclobutyl	propargyl
4243	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	cyclobutyl	propargyl
4244	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclobutyl	propargyl
4245	4-biphenylmethyl	2-thienylmethyl	cyclobutyl	propargyl
4246	pyrimidinylmeth	4-biphenylmethyl	cyclobutyl	propargyl
4247	hyl	pyrimidinylmeth	cyclobutyl	propargyl
4248	2-benzothiazolyl	1		
4249	methyl	2-benzothiazolylme	cyclobutyl	propargyl
4250	2-benzothiophenylmethyl	thyl		
4251	2-thiomethylethyl	2-benzothiophenylmethyl	cyclobutyl	propargyl
4252	1	ethyl		
4253	2-thiomethylmeth	2-thiomethylethyl	cyclobutyl	propargyl
4254	yl			
4255	2-methylpropyl	2-thiomethylmethyl	cyclobutyl	propargyl
4256	2-methylbutyl	2-methylpropyl	cyclobutyl	propargyl
4257	3-methylbutyl	2-methylbutyl	cyclobutyl	propargyl
	cyclopropylmet	3-methylbutyl	cyclobutyl	propargyl
	hyl			
4253	cyclobutylmeth	cyclopropylmeth	cyclobutyl	propargyl
4254	yl	1		
4255	cyclopentylmet	cyclobutylmethyl	cyclobutyl	propargyl
4256	hyl			
4257	p-hydroxyphenyl-methyl	cyclopentylmeth	cyclobutyl	propargyl
		1		
4256	p-nitrophenylmet	p-hydroxyphenyl-methyl	cyclobutyl	propargyl
4257	hyl			
	p-aminophenyl-methyl	p-nitrophenylmeth	cyclobutyl	propargyl
		1		

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4258	"4-(N,N-dimethylamino)phenylmethyl" benzyl	p-aminophenyl-methyl	cyclobutyl	propargyl
4259		"4-(N,N-dimethylamino)phenylmethyl" benzyl	cyclobutylmeth yl	cyclobutyl
4260	1-pyrollylmethyl		cyclobutylmeth yl	cyclobutyl
4261	1-pyrazolylmethy l	1-pyrollylmethyl	cyclobutylmeth yl	cyclobutyl
4262	1-imidazolylmeth yl	1-pyrazolylmethy l	cyclobutylmeth yl	cyclobutyl
4263	1-indolylmethy l	1-imidazolylmethy l	cyclobutylmeth yl	cyclobutyl
4264	1-triazolylmethy l	1-indolylmethy l	cyclobutylmeth yl	cyclobutyl
4265	1-tetrazolylmeth yl	1-triazolylmethy l	cyclobutylmeth yl	cyclobutyl
4266	2-pyridylmethy l	1-tetrazolylmethy l	cyclobutylmeth yl	cyclobutyl
4267	3-pyridylmethy l	2-pyridylmethy l	cyclobutylmeth yl	cyclobutyl
4268	4-pyridylmethy l	3-pyridylmethy l	cyclobutylmeth yl	cyclobutyl
4269	cyclohexylmeth yl	4-pyridylmethy l	cyclobutylmeth yl	cyclobutyl
4270	2-naphthylmethy l	cyclohexylmethy l	cyclobutylmeth yl	cyclobutyl
4271	3-naphthylmethy l	2-naphthylmethy l	cyclobutylmeth yl	cyclobutyl
4272	2-thiophenylmeth yl	3-naphthylmethy l	cyclobutylmeth yl	cyclobutyl
4273	4-(1-methyl)piperid inyl-methyl	2-thiophenylmethy l	cyclobutylmeth yl	cyclobutyl
4274	"(3,4-methylenedioxy phenyl)methyl"	4-(1-methyl)piperidin yl-methyl	cyclobutylmeth yl	cyclobutyl
4275	2-thienylmethyl	"(3,4-methylenedioxyph enyl)methyl"	cyclobutylmeth yl	cyclobutyl
4276	4-biphenylmethyl	2-thienylmethyl	cyclobutylmeth yl	cyclobutyl
4277	pyrimidinylmet hyl	4-biphenylmethyl	cyclobutylmeth yl	cyclobutyl
4278	2-benzothiazolyl methyl	pyrimidinylmethy l	cyclobutylmeth yl	cyclobutyl
4279	2-benzothiopheny lmethyl	2-benzothiazolylme thyl	cyclobutylmeth yl	cyclobutyl

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4280	2-thiomethylethyl	2-benzothiophenylmethyl	cyclobutylmeth yl	cyclobutyl
4281	2-thiomethylmeth yl	2-thiomethylethyl	cyclobutylmeth yl	cyclobutyl
4282	2-methylpropyl	2-thiomethylmeth yl	cyclobutylmeth yl	cyclobutyl
4283	2-methylbutyl	2-methylpropyl	cyclobutylmeth yl	cyclobutyl
4284	3-methylbutyl	2-methylbutyl	cyclobutylmeth yl	cyclobutyl
4285	cyclopropylmeth yl	3-methylbutyl	cyclobutylmeth yl	cyclobutyl
4286	cyclobutylmeth yl	cyclopropylmeth yl	cyclobutylmeth yl	cyclobutyl
4287	cyclopentylmeth yl	cyclobutylmeth yl	cyclobutylmeth yl	cyclobutyl
4288	p-hydroxyphenyl- methyl	cyclopentylmeth yl	cyclobutylmeth yl	cyclobutyl
4289	p-nitrophenylmeth yl	p-hydroxyphenyl- methyl	cyclobutylmeth yl	cyclobutyl
4290	p-aminophenyl- methyl	p-nitrophenylmeth yl	cyclobutylmeth yl	cyclobutyl
4291	"4-(N,N- dimethylamino) phenylmethyl" benzyl	p-aminophenyl- methyl	cyclobutylmeth yl	cyclobutyl
4292		"4-(N,N- dimethylamino)ph enylmethyl" benzyl	cyclopentyl	cyclobutylmeth yl
4293	1-pyrolylmethyl		cyclopentyl	cyclobutylmeth yl
4294	1-pyrazolylmeth yl	1-pyrolylmethyl	cyclopentyl	cyclobutylmeth yl
4295	1-imidazolylmeth yl	1-pyrazolylmeth yl	cyclopentyl	cyclobutylmeth yl
4296	1-indolylmethyl	1-imidazolylmeth yl	cyclopentyl	cyclobutylmeth yl
4297	1-triazolylmeth yl	1-indolylmethyl	cyclopentyl	cyclobutylmeth yl
4298	1-tetrazolylmeth yl	1-triazolylmeth yl	cyclopentyl	cyclobutylmeth yl
4299	2-pyridylmethyl	1-tetrazolylmeth yl	cyclopentyl	cyclobutylmeth yl
4300	3-pyridylmethyl	2-pyridylmethyl	cyclopentyl	cyclobutylmeth yl
4301	4-pyridylmethyl	3-pyridylmethyl	cyclopentyl	cyclobutylmeth yl
4302	cyclohexylmeth yl	4-pyridylmethyl	cyclopentyl	cyclobutylmeth yl

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4303	2-naphthylmethyl	cyclohexylmethyl	cyclopentyl	cyclobutylmethyl
4304	3-naphthylmethyl	2-naphthylmethyl	cyclopentyl	cyclobutylmethyl
4305	2-thiophenylmethyl	3-naphthylmethyl	cyclopentyl	cyclobutylmethyl
4306	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	cyclopentyl	cyclobutylmethyl
4307	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	cyclopentyl	cyclobutylmethyl
4308	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclopentyl	cyclobutylmethyl
4309	4-biphenylmethyl	2-thienylmethyl	cyclopentyl	cyclobutylmethyl
4310	pyrimidinylmethyl	4-biphenylmethyl	cyclopentyl	cyclobutylmethyl
4311	2-benzothiazolylmethyl	pyrimidinylmethyl	cyclopentyl	cyclobutylmethyl
4312	2-benzothiophenylmethyl	2-benzothiazolylmethyl	cyclopentyl	cyclobutylmethyl
4313	2-thiomethylethyl	2-benzothiophenylmethyl	cyclopentyl	cyclobutylmethyl
4314	2-thiomethylmethyl	2-thiomethylethyl	cyclopentyl	cyclobutylmethyl
4315	2-methylpropyl	2-thiomethylmethyl	cyclopentyl	cyclobutylmethyl
4316	2-methylbutyl	2-methylpropyl	cyclopentyl	cyclobutylmethyl
4317	3-methylbutyl	2-methylbutyl	cyclopentyl	cyclobutylmethyl
4318	cyclopropylmethyl	3-methylbutyl	cyclopentyl	cyclobutylmethyl
4319	cyclobutylmethyl	cyclopropylmethyl	cyclopentyl	cyclobutylmethyl
4320	cyclopentylmethyl	cyclobutylmethyl	cyclopentyl	cyclobutylmethyl
4321	p-hydroxyphenylmethyl	cyclopentylmethyl	cyclopentyl	cyclobutylmethyl
4322	p-nitrophenylmethyl	p-hydroxyphenylmethyl	cyclopentyl	cyclobutylmethyl
4323	p-aminophenylmethyl	p-nitrophenylmethyl	cyclopentyl	cyclobutylmethyl
4324	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	cyclopentyl	cyclobutylmethyl

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4325	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	cyclopentylmethyl	cyclopentyl
4326	1-pyrolylmethyl	benzyl	cyclopentylmethyl	cyclopentyl
4327	1-pyrazolylmethyl	1-pyrolylmethyl	cyclopentylmethyl	cyclopentyl
4328	1-imidazolylmethyl	1-pyrazolylmethyl	cyclopentylmethyl	cyclopentyl
4329	1-indolylmethyl	1-imidazolylmethyl	cyclopentylmethyl	cyclopentyl
4330	1-triazolylmethyl	1-indolylmethyl	cyclopentylmethyl	cyclopentyl
4331	1-tetrazolylmethyl	1-triazolylmethyl	cyclopentylmethyl	cyclopentyl
4332	2-pyridylmethyl	1-tetrazolylmethyl	cyclopentylmethyl	cyclopentyl
4333	3-pyridylmethyl	2-pyridylmethyl	cyclopentylmethyl	cyclopentyl
4334	4-pyridylmethyl	3-pyridylmethyl	cyclopentylmethyl	cyclopentyl
4335	cyclohexylmethyl	4-pyridylmethyl	cyclopentylmethyl	cyclopentyl
4336	2-naphthylmethyl	cyclohexylmethyl	cyclopentylmethyl	cyclopentyl
4337	3-naphthylmethyl	2-naphthylmethyl	cyclopentylmethyl	cyclopentyl
4338	2-thiophenylmethyl	3-naphthylmethyl	cyclopentylmethyl	cyclopentyl
4339	4-(1-methyl)piperidinylmethyl	2-thiophenylmethyl	cyclopentylmethyl	cyclopentyl
4340	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinylmethyl	cyclopentylmethyl	cyclopentyl
4341	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclopentylmethyl	cyclopentyl
4342	4-biphenylmethyl	2-thienylmethyl	cyclopentylmethyl	cyclopentyl
4343	pyrimidinylmethyl	4-biphenylmethyl	cyclopentylmethyl	cyclopentyl
4344	2-benzothiazolylmethyl	pyrimidinylmethyl	cyclopentylmethyl	cyclopentyl
4345	2-benzothiophenylmethyl	2-benzothiazolylmethyl	cyclopentylmethyl	cyclopentyl
4346	2-thiomethylethyl	2-benzothiophenylmethyl	cyclopentylmethyl	cyclopentyl

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4347	2-thiomethylmeth yl	2-thiomethylethyl	cyclopentyl- methyl	cyclopentyl
4348	2-methylpropyl	2-thiomethylmethyl	cyclopentyl- methyl	cyclopentyl
4349	2-methylbutyl	2-methylpropyl	cyclopentyl- methyl	cyclopentyl
4350	3-methylbutyl	2-methylbutyl	cyclopentyl- methyl	cyclopentyl
4351	cyclopropylmet hyl	3-methylbutyl	cyclopentyl- methyl	cyclopentyl
4352	cyclobutylmeth yl	cyclopropyl- methyl	cyclopentyl- methyl	cyclopentyl
4353	cyclopentylmet hyl	cyclobutylmethyl	cyclopentyl- methyl	cyclopentyl
4354	p-hydroxyphenyl- methyl	cyclopentyl- methyl	cyclopentyl- methyl	cyclopentyl
4355	p-nitrophenylmet hyl	p-hydroxyphenyl- methyl	cyclopentyl- methyl	cyclopentyl
4356	p-aminophenyl- methyl	p-nitrophenyl- methyl	cyclopentyl- methyl	cyclopentyl
4357	"4-(N,N- dimethylamino) phenylmethyl" benzyl	p-aminophenyl- methyl	cyclopentyl- methyl	cyclopentyl
4358		"4-(N,N- dimethylamino)ph enylmethyl" benzyl	1-hexyl	cyclopentylmeth yl
4359	1-pyrololmethyl		1-hexyl	cyclopentylmeth yl
4360	1-pyrazolylmethy l	1-pyrololmethyl	1-hexyl	cyclopentylmeth yl
4361	1-imidazolylmeth yl	1-pyrazolylmethyl	1-hexyl	cyclopentylmeth yl
4362	1-indolylmethyl	1-imidazolylmethyl	1-hexyl	cyclopentylmeth yl
4363	1-triazolylmethy l	1-indolylmethyl	1-hexyl	cyclopentylmeth yl
4364	1-tetrazolylmeth yl	1-triazolylmethyl	1-hexyl	cyclopentylmeth yl
4365	2-pyridylmethyl	1-tetrazolylmethyl	1-hexyl	cyclopentylmeth yl
4366	3-pyridylmethyl	2-pyridylmethyl	1-hexyl	cyclopentylmeth yl
4367	4-pyridylmethyl	3-pyridylmethyl	1-hexyl	cyclopentylmeth yl
4368	cyclohexylmeth yl	4-pyridylmethyl	1-hexyl	cyclopentylmeth yl
4369	2-naphthylmethyl	cyclohexylmethyl	1-hexyl	cyclopentylmeth yl
4370	3-naphthylmethyl	2-naphthylmethyl	1-hexyl	cyclopentylmeth yl

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4371	2-thiophenylmeth yl	3-naphthylmethyl	1-hexyl	cyclopentylmeth yl
4372	4-(1-methyl)piperid inyl-methyl	2-thiophenylmethyl	1-hexyl	cyclopentylmeth yl
4373	"(3,4-methylenedioxy phenyl)methyl"	4-(1-methyl)piperidin yl-methyl	1-hexyl	cyclopentylmeth yl
4374	2-thienylmethyl	"(3,4-methylenedioxyph enyl)methyl"	1-hexyl	cyclopentylmeth yl
4375	4-biphenylmethyl	2-thienylmethyl	1-hexyl	cyclopentylmeth yl
4376	pyrimidinylmet hyl	4-biphenylmethyl	1-hexyl	cyclopentylmeth yl
4377	2-benzothiazolyl methyl	pyrimidinylmethy l	1-hexyl	cyclopentylmeth yl
4378	2-benzothiopheny lmethyl	2-benzothiazolylme thyl	1-hexyl	cyclopentylmeth yl
4379	2-thiomethylethy l	2-benzothiophenylm ethyl	1-hexyl	cyclopentylmeth yl
4380	2-thiomethylmeth yl	2-thiomethylethyl	1-hexyl	cyclopentylmeth yl
4381	2-methylpropyl	2-thiomethylmethyl	1-hexyl	cyclopentylmeth yl
4382	2-methylbutyl	2-methylpropyl	1-hexyl	cyclopentylmeth yl
4383	3-methylbutyl	2-methylbutyl	1-hexyl	cyclopentylmeth yl
4384	cyclopropylmet hyl	3-methylbutyl	1-hexyl	cyclopentylmeth yl
4385	cyclobutylmeth yl	cyclopropylmethy l	1-hexyl	cyclopentylmeth yl
4386	cyclopentylmet hyl	cyclobutylmethyl	1-hexyl	cyclopentylmeth yl
4387	p-hydroxyphenyl- methyl	cyclopentylmethy l	1-hexyl	cyclopentylmeth yl
4388	p-nitrophenylmet hyl	p-hydroxyphenyl- methyl	1-hexyl	cyclopentylmeth yl
4389	p-aminophenyl- methyl	p-nitrophenylmethy l	1-hexyl	cyclopentylmeth yl
4390	"4-(N,N-dimethylamino) phenylmethyl"	p-aminophenyl- methyl	1-hexyl	cyclopentylmeth yl
4391	benzyl	"4-(N,N-dimethylamino)ph enylmethyl"	4-methylpentyl	1-hexyl
4392	1-pyrolylmethyl	benzyl	4-methylpentyl	1-hexyl

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4393	1-pyrazolylmethyl	1-pyrazolylmethyl	4-methylpentyl	1-hexyl
4394	1-imidazolylmethyl	1-pyrazolylmethyl	4-methylpentyl	1-hexyl
4395	1-indolylmethyl	1-imidazolylmethyl	4-methylpentyl	1-hexyl
4396	1-triazolylmethyl	1-indolylmethyl	4-methylpentyl	1-hexyl
4397	1-tetrazolylmethyl	1-triazolylmethyl	4-methylpentyl	1-hexyl
4398	2-pyridylmethyl	1-tetrazolylmethyl	4-methylpentyl	1-hexyl
4399	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	1-hexyl
4400	4-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	1-hexyl
4401	cyclohexylmethyl	4-pyridylmethyl	4-methylpentyl	1-hexyl
4402	2-naphthylmethyl	cyclohexylmethyl	4-methylpentyl	1-hexyl
4403	3-naphthylmethyl	2-naphthylmethyl	4-methylpentyl	1-hexyl
4404	2-thiophenylmethyl	3-naphthylmethyl	4-methylpentyl	1-hexyl
4405	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	4-methylpentyl	1-hexyl
4406	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	4-methylpentyl	1-hexyl
4407	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	4-methylpentyl	1-hexyl
4408	4-biphenylmethyl	2-thienylmethyl	4-methylpentyl	1-hexyl
4409	pyrimidinylmethyl	4-biphenylmethyl	4-methylpentyl	1-hexyl
4410	2-benzothiazolylmethyl	pyrimidinylmethyl	4-methylpentyl	1-hexyl
4411	2-benzothiophenylmethyl	2-benzothiazolylmethyl	4-methylpentyl	1-hexyl
4412	2-thiomethylethyl	2-benzothiophenylmethyl	4-methylpentyl	1-hexyl
4413	2-thiomethylmethyl	2-thiomethylethyl	4-methylpentyl	1-hexyl
4414	2-methylpropyl	2-thiomethylmethyl	4-methylpentyl	1-hexyl
4415	2-methylbutyl	2-methylpropyl	4-methylpentyl	1-hexyl

Table 6

4416	3-methylbutyl	2-methylbutyl	4-methylpentyl	1-hexyl
4417	cyclopropylmeth yl	3-methylbutyl	4-methylpentyl	1-hexyl
4418	cyclobutylmeth yl	cyclopropylmethy l	4-methylpentyl	1-hexyl
4419	cyclopentylmeth yl	cyclobutylmethy l	4-methylpentyl	1-hexyl
4420	p- hydroxyphenyl- methyl	cyclopentylmethy l	4-methylpentyl	1-hexyl
4421	p- nitrophenylmeth yl	p-hydroxyphenyl- methyl	4-methylpentyl	1-hexyl
4422	p-aminophenyl- methyl	p-nitrophenyl- methyl	4-methylpentyl	1-hexyl
4423	"4-(N,N- dimethylamino) phenylmethyl" benzyl	p-aminophenyl- methyl	4-methylpentyl	1-hexyl
4424		"4-(N,N- dimethyl-amino)- phenylmethyl" benzyl	3-methylpentyl	4-methylpentyl
4425	1-pyrrolyl- methyl		3-methylpentyl	4-methylpentyl
4426	1-pyrazolyl- methyl	1-pyrrolylmethyl	3-methylpentyl	4-methylpentyl
4427	1-imidazolyl- methyl	1- pyrazolylmethyl	3-methylpentyl	4-methylpentyl
4428	1-indolyl- methyl	1- imidazolylmethyl	3-methylpentyl	4-methylpentyl
4429	1-triazolyl- methyl	1-indolylmethyl	3-methylpentyl	4-methylpentyl
4430	1-tetrazolyl- methyl	1- triazolylmethyl	3-methylpentyl	4-methylpentyl
4431	2-pyridyl- methyl	1- tetrazolylmethyl	3-methylpentyl	4-methylpentyl
4432	3- pyridylmethyl	2-pyridylmethyl	3-methylpentyl	4-methylpentyl
4433	4- pyridylmethyl	3-pyridylmethyl	3-methylpentyl	4-methylpentyl
4434	cyclohexylmeth yl	4-pyridylmethyl	3-methylpentyl	4-methylpentyl
4435	2- naphthylmethyl	cyclohexylmethyl	3-methylpentyl	4-methylpentyl
4436	3- naphthylmethyl	2-naphthylmethyl	3-methylpentyl	4-methylpentyl
4437	2-thiophenyl- methyl	3-naphthylmethyl	3-methylpentyl	4-methylpentyl
4438	4-(1- methyl)piperid inyl-methyl	2- thiophenylmethyl	3-methylpentyl	4-methylpentyl
4439	"(3,4- methylenedioxy phenyl)methyl"	4-(1- methyl)piperidin yl-methyl	3-methylpentyl	4-methylpentyl
4440	2- thienylmethyl	"(3,4- methylenedioxyph enyl)methyl"	3-methylpentyl	4-methylpentyl

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4441	4-biphenylmethyl	2-thienylmethyl	3-methylpentyl	4-methylpentyl
4442	pyrimidinylmethyl	4-biphenylmethyl	3-methylpentyl	4-methylpentyl
4443	2-benzothiazolylmethyl	pyrimidinylmethyl	3-methylpentyl	4-methylpentyl
4444	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-methylpentyl	4-methylpentyl
4445	2-thiomethylethyl	2-benzothiophenylmethyl	3-methylpentyl	4-methylpentyl
4446	2-thiomethylmethyl	2-thiomethylethyl	3-methylpentyl	4-methylpentyl
4447	2-methylpropyl	2-thiomethylmethyl	3-methylpentyl	4-methylpentyl
4448	2-methylbutyl	2-methylpropyl	3-methylpentyl	4-methylpentyl
4449	3-methylbutyl	2-methylbutyl	3-methylpentyl	4-methylpentyl
4450	cyclopropylmethyl	3-methylbutyl	3-methylpentyl	4-methylpentyl
4451	cyclobutylmethyl	cyclopropylmethyl	3-methylpentyl	4-methylpentyl
4452	cyclopentylmethyl	cyclobutylmethyl	3-methylpentyl	4-methylpentyl
4453	p-hydroxyphenylmethyl	cyclopentylmethyl	3-methylpentyl	4-methylpentyl
4454	p-nitrophenylmethyl	p-hydroxyphenylmethyl	3-methylpentyl	4-methylpentyl
4455	p-aminophenylmethyl	p-nitrophenylmethyl	3-methylpentyl	4-methylpentyl
4456	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	3-methylpentyl	4-methylpentyl
4457	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	isopropyl	3-methylpentyl
4458	1-pyrollylmethyl	benzyl	isopropyl	3-methylpentyl
4459	1-pyrazolylmethyl	1-pyrollylmethyl	isopropyl	3-methylpentyl
4460	1-imidazolylmethyl	1-pyrazolylmethyl	isopropyl	3-methylpentyl
4461	1-indolylmethyl	1-imidazolylmethyl	isopropyl	3-methylpentyl
4462	1-triazolylmethyl	1-indolylmethyl	isopropyl	3-methylpentyl
4463	1-tetrazolylmethyl	1-triazolylmethyl	isopropyl	3-methylpentyl

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4464	2-pyridylmethyl	1-tetrazolylmethyl	isopropyl	3-methylpentyl
4465	3-pyridylmethyl	2-pyridylmethyl	isopropyl	3-methylpentyl
4466	4-pyridylmethyl	3-pyridylmethyl	isopropyl	3-methylpentyl
4467	cyclohexylmethyl	4-pyridylmethyl	isopropyl	3-methylpentyl
4468	2-naphthylmethyl	cyclohexylmethyl	isopropyl	3-methylpentyl
4469	3-naphthylmethyl	2-naphthylmethyl	isopropyl	3-methylpentyl
4470	2-thiophenylmethyl	3-naphthylmethyl	isopropyl	3-methylpentyl
4471	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	isopropyl	3-methylpentyl
4472	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	isopropyl	3-methylpentyl
4473	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	isopropyl	3-methylpentyl
4474	4-biphenylmethyl	2-thienylmethyl	isopropyl	3-methylpentyl
4475	pyrimidinylmethyl	4-biphenylmethyl	isopropyl	3-methylpentyl
4476	2-benzothiazolylmethyl	pyrimidinylmethyl	isopropyl	3-methylpentyl
4477	2-benzothio-phenylmethyl	2-benzothiazolylmethyl	isopropyl	3-methylpentyl
4478	2-thiomethyl-ethyl	2-benzothiophenylmethyl	isopropyl	3-methylpentyl
4479	2-thiomethyl-methyl	2-thiomethylethyl	isopropyl	3-methylpentyl
4480	2-methylpropyl	2-thiomethylmethyl	isopropyl	3-methylpentyl
4481	2-methylbutyl	2-methylpropyl	isopropyl	3-methylpentyl
4482	3-methylbutyl	2-methylbutyl	isopropyl	3-methylpentyl
4483	cyclopropyl-methyl	3-methylbutyl	isopropyl	3-methylpentyl
4484	cyclobutyl-methyl	cyclopropyl-methyl	isopropyl	3-methylpentyl
4485	cyclopentyl-methyl	cyclobutylmethyl	isopropyl	3-methylpentyl
4486	p-hydroxyphenyl-methyl	cyclopentyl-methyl	isopropyl	3-methylpentyl
4487	p-nitrophenyl-methyl	p-hydroxyphenyl-methyl	isopropyl	3-methylpentyl
4488	p-aminophenyl-methyl	p-nitrophenyl-methyl	isopropyl	3-methylpentyl

Table 6

4489	"4-(N,N-dimethylamino)phenylmethyl" benzyl	p-aminophenyl-methyl	isopropyl	3-methylpentyl
4490		"4-(N,N-dimethylamino)phenylmethyl" benzyl	3-methylbutyl	isopropyl
4491	1-pyrolylmethyl		3-methylbutyl	isopropyl
4492	1-pyrazolyl-methyl	1-pyrolylmethyl	3-methylbutyl	isopropyl
4493	1-imidazolylmeth	1-pyrazolylmethyl	3-methylbutyl	isopropyl
4494	1-indolylmethyl	1-imidazolylmethyl	3-methylbutyl	isopropyl
4495	1-triazolylmethy	1-indolylmethyl	3-methylbutyl	isopropyl
4496	1-tetrazolylmeth	1-triazolylmethyl	3-methylbutyl	isopropyl
4497	2-pyridylmethyl	1-tetrazolylmethyl	3-methylbutyl	isopropyl
4498	3-pyridylmethyl	2-pyridylmethyl	3-methylbutyl	isopropyl
4499	4-pyridylmethyl	3-pyridylmethyl	3-methylbutyl	isopropyl
4500	cyclohexylmeth	4-pyridylmethyl	3-methylbutyl	isopropyl
4501	2-naphthylmethyl	cyclohexylmethyl	3-methylbutyl	isopropyl
4502	3-naphthylmethyl	2-naphthylmethyl	3-methylbutyl	isopropyl
4503	2-thiophenylmeth	3-naphthylmethyl	3-methylbutyl	isopropyl
4504	4-(1-methyl)piperid	2-thiophenylmethyl	3-methylbutyl	isopropyl
4505	inyl-methyl	4-(1-methyl)piperidin	3-methylbutyl	isopropyl
4506	"(3,4-methylenedioxyphenyl)methyl"	yl-methyl	3-methylbutyl	isopropyl
4507	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-methylbutyl	isopropyl
4508	4-biphenylmethyl	2-thienylmethyl	3-methylbutyl	isopropyl
4509	pyrimidinylmet	4-biphenylmethyl	3-methylbutyl	isopropyl
4510	2-benzothiazolyl	pyrimidinylmethy	3-methylbutyl	isopropyl
	1-methyl	1		
	2-benzothiopheny	2-benzothiazolylme	3-methylbutyl	isopropyl
	lmethyl	thyl		

Table 6

4511	2-thiomethylethyl	2-benzothiophenylmethyl	3-methylbutyl	isopropyl
4512	1-thiomethylmethylethyl	2-thiomethylethyl	3-methylbutyl	isopropyl
4513	2-methylpropyl	2-thiomethylmethylethyl	3-methylbutyl	isopropyl
4514	2-methylbutyl	2-methylpropyl	3-methylbutyl	isopropyl
4515	3-methylbutyl	2-methylbutyl	3-methylbutyl	isopropyl
4516	cyclopropylmethyl	3-methylbutyl	3-methylbutyl	isopropyl
4517	cyclobutylmethyl	cyclopropylmethyl	3-methylbutyl	isopropyl
4518	cyclopentylmethyl	cyclobutylmethyl	3-methylbutyl	isopropyl
4519	p-hydroxyphenylmethyl	cyclopentylmethyl	3-methylbutyl	isopropyl
4520	p-nitrophenylmethyl	p-hydroxyphenylmethyl	3-methylbutyl	isopropyl
4521	p-aminophenylmethyl	p-nitrophenylmethyl	3-methylbutyl	isopropyl
4522	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	3-methylbutyl	isopropyl
4523	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-phenylethyl	3-methylbutyl
4524	1-pyrollylmethyl	benzyl	2-phenylethyl	3-methylbutyl
4525	1-pyrazolylmethylethyl	1-pyrollylmethyl	2-phenylethyl	3-methylbutyl
4526	1-imidazolylmethyl	1-pyrazolylmethylethyl	2-phenylethyl	3-methylbutyl
4527	1-indolylmethyl	1-imidazolylmethyl	2-phenylethyl	3-methylbutyl
4528	1-triazolylmethylethyl	1-indolylmethyl	2-phenylethyl	3-methylbutyl
4529	1-tetrazolylmethylethyl	1-triazolylmethylethyl	2-phenylethyl	3-methylbutyl
4530	2-pyridylmethyl	1-tetrazolylmethylethyl	2-phenylethyl	3-methylbutyl
4531	3-pyridylmethyl	2-pyridylmethyl	2-phenylethyl	3-methylbutyl
4532	4-pyridylmethyl	3-pyridylmethyl	2-phenylethyl	3-methylbutyl
4533	cyclohexylmethyl	4-pyridylmethyl	2-phenylethyl	3-methylbutyl

Table 6

4534	2-naphthylmethyl	cyclohexylmethyl	2-phenylethyl	3-methylbutyl
4535	3-naphthylmethyl	2-naphthylmethyl	2-phenylethyl	3-methylbutyl
4536	2-thiophenylmethyl	3-naphthylmethyl	2-phenylethyl	3-methylbutyl
4537	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-phenylethyl	3-methylbutyl
4538	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-phenylethyl	3-methylbutyl
4539	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-phenylethyl	3-methylbutyl
4540	4-biphenylmethyl	2-thienylmethyl	2-phenylethyl	3-methylbutyl
4541	pyrimidinylmethyl	4-biphenylmethyl	2-phenylethyl	3-methylbutyl
4542	2-benzothiazolylmethyl	pyrimidinylmethyl	2-phenylethyl	3-methylbutyl
4543	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-phenylethyl	3-methylbutyl
4544	2-thiomethylethyl	2-benzothiophenylmethyl	2-phenylethyl	3-methylbutyl
4545	2-thiomethylmethyl	2-thiomethylethyl	2-phenylethyl	3-methylbutyl
4546	2-methylpropyl	2-thiomethylmethyl	2-phenylethyl	3-methylbutyl
4547	2-methylbutyl	2-methylpropyl	2-phenylethyl	3-methylbutyl
4548	3-methylbutyl	2-methylbutyl	2-phenylethyl	3-methylbutyl
4549	cyclopropylmethyl	3-methylbutyl	2-phenylethyl	3-methylbutyl
4550	cyclobutylmethyl	cyclopropylmethyl	2-phenylethyl	3-methylbutyl
4551	cyclopentylmethyl	cyclobutylmethyl	2-phenylethyl	3-methylbutyl
4552	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-phenylethyl	3-methylbutyl
4553	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-phenylethyl	3-methylbutyl

Table 6

4554	p-aminophenyl-methyl	p-nitrophenylmethyl	2-phenylethyl	3-methylbutyl
4555	"4-(N,N-dimethylamino)phenylmethyl	p-aminophenyl-methyl	2-phenylethyl	3-methylbutyl
4556	benzyl	"4-(N,N-dimethylamino)phenylmethyl	3-phenylpropyl	2-phenylethyl
4557	1-pyrololymethyl	benzyl	3-phenylpropyl	2-phenylethyl
4558	1-pyrazololymethyl	1-pyrololymethyl	3-phenylpropyl	2-phenylethyl
4559	1-imidazololymethyl	1-pyrazololymethyl	3-phenylpropyl	2-phenylethyl
4560	1-indololymethyl	1-imidazololymethyl	3-phenylpropyl	2-phenylethyl
4561	1-triazololymethyl	1-indololymethyl	3-phenylpropyl	2-phenylethyl
4562	1-tetrazololymethyl	1-triazololymethyl	3-phenylpropyl	2-phenylethyl
4563	2-pyridololymethyl	1-tetrazololymethyl	3-phenylpropyl	2-phenylethyl
4564	3-pyridololymethyl	2-pyridololymethyl	3-phenylpropyl	2-phenylethyl
4565	4-pyridololymethyl	3-pyridololymethyl	3-phenylpropyl	2-phenylethyl
4566	cyclohexololymethyl	4-pyridololymethyl	3-phenylpropyl	2-phenylethyl
4567	2-naphthololymethyl	1-cyclohexololymethyl	3-phenylpropyl	2-phenylethyl
4568	3-naphthololymethyl	2-naphthololymethyl	3-phenylpropyl	2-phenylethyl
4569	1-thiophenololymethyl	1-naphthololymethyl	3-phenylpropyl	2-phenylethyl
4570	4-(1-methyl)piperidinololymethyl	2-thiophenololymethyl	3-phenylpropyl	2-phenylethyl
4571	"(3,4-methylenedioxyphenyl)methyl	1-methylpiperidinololymethyl	3-phenylpropyl	2-phenylethyl
4572	2-thienololymethyl	"(3,4-methylenedioxyphenyl)methyl	3-phenylpropyl	2-phenylethyl

Table 6

4573	4-biphenylmethyl	2-thienylmethyl	3-phenylpropyl	2-phenylethyl
4574	1-pyrimidinylmethyl	4-biphenylmethyl	3-phenylpropyl	2-phenylethyl
4575	2-benzothiazolylmethyl	1-pyrimidinylmethyl	3-phenylpropyl	2-phenylethyl
4576	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-phenylpropyl	2-phenylethyl
4577	2-thiomethylethyl	2-benzothiophenylmethyl	3-phenylpropyl	2-phenylethyl
4578	2-thiomethylmethyl	2-thiomethylethyl	3-phenylpropyl	2-phenylethyl
4579	2-methylpropyl	2-thiomethylmethyl	3-phenylpropyl	2-phenylethyl
4580	2-methylbutyl	2-methylpropyl	3-phenylpropyl	2-phenylethyl
4581	3-methylbutyl	2-methylbutyl	3-phenylpropyl	2-phenylethyl
4582	cyclopropylmethyl	3-methylbutyl	3-phenylpropyl	2-phenylethyl
4583	cyclobutylmethyl	cyclopropylmethyl	3-phenylpropyl	2-phenylethyl
4584	cyclopentylmethyl	cyclobutylmethyl	3-phenylpropyl	2-phenylethyl
4585	p-hydroxyphenylmethyl	cyclopentylmethyl	3-phenylpropyl	2-phenylethyl
4586	p-nitrophenylmethyl	p-hydroxyphenylmethyl	3-phenylpropyl	2-phenylethyl
4587	p-aminophenylmethyl	p-nitrophenylmethyl	3-phenylpropyl	2-phenylethyl
4588	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	3-phenylpropyl	2-phenylethyl
4589	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4590	1-pyrollylmethyl	benzyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4591	1-pyrazolylmethyl	1-pyrollylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4592	1-imidazolylmethyl	1-pyrazolylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl

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4593	1-indolylmethyl	1-imidazolylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4594	1-triazolylmethyl	1-indolylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4595	1-tetrazolylmethyl	1-triazolylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4596	2-pyridylmethyl	1-tetrazolylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4597	3-pyridylmethyl	2-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4598	4-pyridylmethyl	3-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4599	cyclohexylmethyl	4-pyridylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4600	2-naphthylmethyl	cyclohexylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4601	3-naphthylmethyl	2-naphthylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4602	2-thiophenylmethyl	1-naphthylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4603	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4604	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4605	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4606	4-biphenylmethyl	2-thienylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4607	1-pyrimidinylmethyl	4-biphenylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4608	2-benzothiazolylmethyl	1-pyrimidinylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4609	2-benzothiophenylmethyl	2-benzothiazolylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl
4610	2-thiomethylethyl	2-benzothiophenylmethyl	"2-(N,N-dimethylamino)ethyl"	3-phenylpropyl

Table 6

4611	2-thiomethylmet hyl	2-thiomethyleth yl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4612	2-methylpropyl	2-thiomethylmet hyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4613	2-methylbutyl	2-methylpropyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4614	3-methylbutyl	2-methylbutyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4615	cyclopropylmet hyl	3-methylbutyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4616	cyclobutylmet hyl	cyclopropylmet hyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4617	cyclopentylmet hyl	cyclobutylmet hyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4618	p-hydroxyphenyl -methyl	cyclopentylmet hyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4619	p-nitrophenylmet hyl	p-hydroxyphenyl -methyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4620	p-aminophenyl- methyl	p-nitrophenylmet hyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4621	"4-(N,N- dimethylamino)phenylmethyl "	p-aminophenyl- methyl	"2-(N,N- dimethylamino)ethyl"	3-phenylpropyl
4622	benzyl	"4-(N,N- dimethylamino)phenylmethyl "	3-oxetanyl- methyl	"2-(N,N- dimethyl- amino)ethyl"
4623	1-pyrololmethyl	benzyl	3-oxetanyl- methyl	"2-(N,N- dimethyl- amino)ethyl"
4624	1-pyrazolyl- methyl	1-pyrolol- methyl	3-oxetanyl- methyl	"2-(N,N- dimethyl- amino)ethyl"
4625	1-imidazolyl- methyl	1-pyrazolylmeth yl	3-oxetanyl- methyl	"2-(N,N- dimethylamino)ethyl"
4626	1-indolylmethyl	1-imidazolylmet hyl	3-oxetanyl- methyl	"2-(N,N- dimethylamino)ethyl"
4627	1-triazolyl- methyl	1-indolylmethyl	3-oxetanyl- methyl	"2-(N,N- dimethylamino)ethyl"
4628	1-tetrazolylmet hyl	1-triazolyl- methyl	3-oxetanyl- methyl	"2-(N,N- dimethylamino)ethyl"

Table 6

4629	2-pyridylmethyl	1-tetrazolyl-methyl	3-oxetanyl-methyl	"2-(N,N-dimethylamino)ethyl"
4630	3-pyridylmethyl	2-pyridylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4631	4-pyridylmethyl	3-pyridylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4632	cyclohexylmethyl	4-pyridylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4633	2-naphthylmethyl	cyclohexylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4634	3-naphthylmethyl	2-naphthylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4635	2-thiophenylmethyl	3-naphthylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4636	4-(1-methyl)piperidinylmethyl	2-thiophenylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4637	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4638	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4639	4-biphenylmethyl	2-thienylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4640	pyrimidinylmethyl	4-biphenylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4641	2-benzothiazolylmethyl	pyrimidinylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4642	2-benzothiophenylmethyl	2-benzothiazolylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4643	2-thiomethylethyl	2-benzothiophenylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4644	2-thiomethylmethyl	2-thiomethylethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4645	2-methylpropyl	2-thiomethylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4646	2-methylbutyl	2-methylpropyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"

Table 6

4647	3-methylbutyl	2-methylbutyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4648	cyclopropylmethyl	3-methylbutyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4649	cyclobutylmethyl	cyclopropylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4650	cyclopentylmethyl	cyclobutylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4651	p-hydroxyphenylmethyl	cyclopentylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4652	p-nitrophenylmethyl	p-hydroxyphenylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4653	p-aminophenylmethyl	p-nitrophenylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4654	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	3-oxetanylmethyl	"2-(N,N-dimethylamino)ethyl"
4655	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-tetrahydrofuran	3-oxetanylmethyl
4656	1-pyrollylmethyl	benzyl	2-tetrahydrofuran	3-oxetanylmethyl
4657	1-pyrazolylmethyl	1-pyrollylmethyl	2-tetrahydrofuran	3-oxetanylmethyl
4658	1-imidazolylmethyl	1-pyrazolylmethyl	2-tetrahydrofuran	3-oxetanylmethyl
4659	1-indolylmethyl	1-imidazolylmethyl	2-tetrahydrofuran	3-oxetanylmethyl
4660	1-triazolylmethyl	1-indolylmethyl	2-tetrahydrofuran	3-oxetanylmethyl
4661	1-tetrazolylmethyl	1-triazolylmethyl	2-tetrahydrofuran	3-oxetanylmethyl
4662	2-pyridylmethyl	1-tetrazolylmethyl	2-tetrahydrofuran	3-oxetanylmethyl
4663	3-pyridylmethyl	2-pyridylmethyl	2-tetrahydrofuran	3-oxetanylmethyl
4664	4-pyridylmethyl	3-pyridylmethyl	2-tetrahydrofuran	3-oxetanylmethyl

Table 6

4665	cyclohexylmet hyl	4- pyridylmethyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4666	2- naphthylmethy 1	cyclohexylmet hyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4667	3- naphthylmethy 1	2- naphthylmethy 1	2- tetrahydrofur anyl	3- oxetanylmethy 1
4668	2- thiophenylmet hyl	3- naphthylmethy 1	2- tetrahydrofur anyl	3- oxetanylmethy 1
4669	4-(1- methyl)piperi diny1-methyl	2- thiophenylmet hyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4670	"(3,4- methylenediox yphenyl)methy 1"	4-(1- methyl)piperi diny1-methyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4671	2- thienylmethyl	"(3,4- methylenediox yphenyl)methy 1"	2- tetrahydrofur anyl	3- oxetanylmethy 1
4672	4- biphenylmethy 1	2- thienylmethyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4673	pyrimidinylme thyl	4- biphenylmethy 1	2- tetrahydrofur anyl	3- oxetanylmethy 1
4674	2- benzothiazoly lmethyl	pyrimidinylme thyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4675	2- benzothiophen ylmethyl	2- benzothiazoly lmethyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4676	2- thiomethyleth yl	2- benzothiophen ylmethyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4677	2- thiomethylmet hyl	2- thiomethyleth yl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4678	2- methylpropyl	2- thiomethylmet hyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4679	2-methylbutyl	2- methylpropyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4680	3-methylbutyl	2-methylbutyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4681	cyclopropylme thyl	3-methylbutyl	2- tetrahydrofur anyl	3- oxetanylmethy 1
4682	cyclobutylmet hyl	cyclopropylme thyl	2- tetrahydrofur anyl	3- oxetanylmethy 1

Table 6

4683	cyclopentylmeth thyl	cyclobutylmeth hyl	2- tetrahydrofur anyl	3- oxetanylmethy l
4684	p- hydroxyphenyl -methyl	cyclopentylmeth thyl	2- tetrahydrofur anyl	3- oxetanylmethy l
4685	p- nitrophenylmeth thyl	p- hydroxyphenyl -methyl	2- tetrahydrofur anyl	3- oxetanylmethy l
4686	p- aminophenyl- methyl	p- nitrophenylmeth thyl	2- tetrahydrofur anyl	3- oxetanylmethy l
4687	"4-(N,N- dimethylamino)phenylmethyl "	p- aminophenyl- methyl	2- tetrahydrofur anyl	3- oxetanylmethy l
4688	benzyl	"4-(N,N- dimethylamino)phenylmethyl "	2- methoxypropyl	2- tetrahydrofur anyl
4689	1- pyrolylmethyl	benzyl	2- methoxypropyl	2- tetrahydrofur anyl
4690	1- pyrazolylmeth yl	1- pyrolylmethyl	2- methoxypropyl	2- tetrahydrofur anyl
4691	1- imidazolylmeth hyl	1- pyrazolylmeth yl	2- methoxypropyl	2- tetrahydrofur anyl
4692	1- indolylmethyl	1- imidazolylmeth hyl	2- methoxypropyl	2- tetrahydrofur anyl
4693	1- triazolylmeth yl	1- indolylmethyl	2- methoxypropyl	2- tetrahydrofur anyl
4694	1- tetrazolylmeth hyl	1- triazolylmeth yl	2- methoxypropyl	2- tetrahydrofur anyl
4695	2- pyridylmethyl	1- tetrazolylmeth hyl	2- methoxypropyl	2- tetrahydrofur anyl
4696	3- pyridylmethyl	2- pyridylmethyl	2- methoxypropyl	2- tetrahydrofur anyl
4697	4- pyridylmethyl	3- pyridylmethyl	2- methoxypropyl	2- tetrahydrofur anyl
4698	cyclohexylmeth hyl	4- pyridylmethyl	2- methoxypropyl	2- tetrahydrofur anyl
4699	2- naphthylmethy l	cyclohexylmeth hyl	2- methoxypropyl	2- tetrahydrofur anyl
4700	3- naphthylmethy l	2- naphthylmethy l	2- methoxypropyl	2- tetrahydrofur anyl

Table 6

4701	2-thiophenylmethyl	3-naphthylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4702	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4703	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl-methyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4704	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-methoxypropyl	2-tetrahydrofuran-2-yl
4705	4-biphenylmethyl	2-thienylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4706	1-pyrimidinylmethyl	4-biphenylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4707	2-benzothiazolylmethyl	1-pyrimidinylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4708	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4709	2-thiomethylethyl	2-benzothiophenylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4710	2-thiomethylmethyl	2-thiomethylethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4711	2-methylpropyl	2-thiomethylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4712	2-methylbutyl	2-methylpropyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4713	3-methylbutyl	2-methylbutyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4714	cyclopropylmethyl	3-methylbutyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4715	cyclobutylmethyl	cyclopropylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4716	cyclopentylmethyl	cyclobutylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4717	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-methoxypropyl	2-tetrahydrofuran-2-yl
4718	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-methoxypropyl	2-tetrahydrofuran-2-yl

Table 6

4719	p-aminophenyl-methyl	p-nitrophenylmethyl	2-methoxypropyl	2-tetrahydrofuran
4720	"4-(N,N-dimethylamino)phenylmethyl	p-aminophenyl-methyl	2-methoxypropyl	2-tetrahydrofuran
4721	benzyl	"4-(N,N-dimethylamino)phenylmethyl	2-ethoxyethyl	2-methoxypropyl
4722	1-pyrolylmethyl	benzyl	2-ethoxyethyl	2-methoxypropyl
4723	1-pyrazolylmethyl	1-pyrolylmethyl	2-ethoxyethyl	2-methoxypropyl
4724	1-imidazolylmethyl	1-pyrazolylmethyl	2-ethoxyethyl	2-methoxypropyl
4725	1-indolylmethyl	1-imidazolylmethyl	2-ethoxyethyl	2-methoxypropyl
4726	1-triazolylmethyl	1-indolylmethyl	2-ethoxyethyl	2-methoxypropyl
4727	1-tetrazolylmethyl	1-triazolylmethyl	2-ethoxyethyl	2-methoxypropyl
4728	2-pyridylmethyl	1-tetrazolylmethyl	2-ethoxyethyl	2-methoxypropyl
4729	3-pyridylmethyl	2-pyridylmethyl	2-ethoxyethyl	2-methoxypropyl
4730	4-pyridylmethyl	3-pyridylmethyl	2-ethoxyethyl	2-methoxypropyl
4731	cyclohexylmethyl	4-pyridylmethyl	2-ethoxyethyl	2-methoxypropyl
4732	2-naphthylmethyl	cyclohexylmethyl	2-ethoxyethyl	2-methoxypropyl
4733	3-naphthylmethyl	2-naphthylmethyl	2-ethoxyethyl	2-methoxypropyl
4734	2-thiophenylmethyl	3-naphthylmethyl	2-ethoxyethyl	2-methoxypropyl
4735	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-ethoxyethyl	2-methoxypropyl
4736	"(3,4-methylenedioxyphenyl)methyl	4-(1-methyl)piperidinyl-methyl	2-ethoxyethyl	2-methoxypropyl
4737	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl	2-ethoxyethyl	2-methoxypropyl

Table 6

4738	4-biphenylmethyl	2-thienylmethyl	2-ethoxyethyl	2-methoxypropyl
4739	pyrimidinylmethyl	4-biphenylmethyl	2-ethoxyethyl	2-methoxypropyl
4740	2-benzothiazolylmethyl	pyrimidinylmethyl	2-ethoxyethyl	2-methoxypropyl
4741	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-ethoxyethyl	2-methoxypropyl
4742	2-thiomethylethyl	2-benzothiophenylmethyl	2-ethoxyethyl	2-methoxypropyl
4743	2-thiomethylmethyl	2-thiomethylethyl	2-ethoxyethyl	2-methoxypropyl
4744	2-methylpropyl	2-thiomethylmethyl	2-ethoxyethyl	2-methoxypropyl
4745	2-methylbutyl	2-methylpropyl	2-ethoxyethyl	2-methoxypropyl
4746	3-methylbutyl	2-methylbutyl	2-ethoxyethyl	2-methoxypropyl
4747	cyclopropylmethyl	3-methylbutyl	2-ethoxyethyl	2-methoxypropyl
4748	cyclobutylmethyl	cyclopropylmethyl	2-ethoxyethyl	2-methoxypropyl
4749	cyclopentylmethyl	cyclobutylmethyl	2-ethoxyethyl	2-methoxypropyl
4750	p-hydroxyphenylmethyl	cyclopentylmethyl	2-ethoxyethyl	2-methoxypropyl
4751	p-nitrophenylmethyl	p-hydroxyphenylmethyl	2-ethoxyethyl	2-methoxypropyl
4752	p-aminophenylmethyl	p-nitrophenylmethyl	2-ethoxyethyl	2-methoxypropyl
4753	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	2-ethoxyethyl	2-methoxypropyl
4754	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-(1-pyrol)ethyl	2-ethoxyethyl
4755	1-pyrolmethyl	benzyl	2-(1-pyrol)ethyl	2-ethoxyethyl
4756	1-pyrazolylmethyl	1-pyrolmethyl	2-(1-pyrol)ethyl	2-ethoxyethyl
4757	1-imidazolylmethyl	1-pyrazolylmethyl	2-(1-pyrol)ethyl	2-ethoxyethyl

Table 6

4758	1-indolylmethyl	1-imidazolylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4759	1-triazolylmethyl	1-indolylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4760	1-tetrazolylmethyl	1-triazolylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4761	2-pyridylmethyl	1-tetrazolylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4762	3-pyridylmethyl	2-pyridylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4763	4-pyridylmethyl	3-pyridylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4764	1-cyclohexylmethyl	4-pyridylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4765	2-naphthylmethyl	1-cyclohexylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4766	3-naphthylmethyl	2-naphthylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4767	2-thiophenylmethyl	3-naphthylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4768	4-(1-methyl)piperidinyl	2-thiophenylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4769	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4770	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4771	4-biphenylmethyl	2-thienylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4772	1-pyrimidinylmethyl	4-biphenylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4773	2-benzothiazolylmethyl	1-pyrimidinylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4774	2-benzothienylmethyl	2-benzothiazolylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4775	2-thiomethyl-ethyl	2-benzothienylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4776	2-thiomethyl-methyl	2-thiomethyl-ethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4777	2-methylpropyl	2-thiomethyl-methyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl

Table 6

4778	2-methylbutyl	2-methylpropyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4779	3-methylbutyl	2-methylbutyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4780	cyclopropyl-methyl	3-methylbutyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4781	cyclobutyl-methyl	cyclopropylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4782	cyclopentyl-methyl	cyclobutylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4783	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4784	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4785	p-aminophenyl-methyl	p-nitrophenylmethyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4786	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-(1-pyrolyl)ethyl	2-ethoxyethyl
4787	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4788	1-pyrolylmethyl	benzyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4789	1-pyrazolylmethyl	1-pyrolylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4790	1-imidazolylmethyl	1-pyrazolylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4791	1-indolylmethyl	1-imidazolylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4792	1-triazolylmethyl	1-indolylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4793	1-tetrazolylmethyl	1-triazolylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4794	2-pyridylmethyl	1-tetrazolylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4795	3-pyridylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4796	4-pyridylmethyl	3-pyridylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl
4797	cyclohexylmethyl	4-pyridylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrolyl)ethyl

Table 6

4798	2-naphthylmethy 1	cyclohexylmet hyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4799	3-naphthylmethy 1	2-naphthylmethy 1	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4800	2-thiophenylmet hyl	3-naphthylmethy 1	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4801	4-(1-methyl)piperi diny1-methyl	2-thiophenylmet hyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4802	"(3,4-methylenedio yphenyl)methy 1"	4-(1-methyl)piperi diny1-methyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4803	2-thienylmethy1	"(3,4-methylenedio yphenyl)methy 1"	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4804	4-biphenylmethy 1	2-thienylmethy1	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4805	pyrimidinylme thyl	4-biphenylmethy 1	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4806	2-benzothiazoly lmethyl	pyrimidinylme thyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4807	2-benzothiophen ylmethy1	2-benzothiazoly lmethyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4808	2-thiomethyleth yl	2-benzothiophen ylmethy1	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4809	2-thiomethylmet hyl	2-thiomethyleth yl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4810	2-methylpropyl	2-thiomethylmet hyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4811	2-methylbutyl	2-methylpropyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4812	3-methylbutyl	2-methylbutyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4813	cyclopropylme thyl	3-methylbutyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4814	cyclobutylmet hyl	cyclopropylme thyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl
4815	cyclopentylme thyl	cyclobutylmet hyl	2-(1-imidazolyl)et hyl	2-(1-pyrolyl)ethyl

Table 6

4816	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrrolyl)ethyl
4817	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-(1-imidazolyl)ethyl	2-(1-pyrrolyl)ethyl
4818	p-aminophenyl-methyl	p-nitrophenylmethyl	2-(1-imidazolyl)ethyl	2-(1-pyrrolyl)ethyl
4819	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenyl-methyl	2-(1-imidazolyl)ethyl	2-(1-pyrrolyl)ethyl
4820	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4821	1-pyrrolylmethyl	benzyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4822	1-pyrazolylmethyl	1-pyrrolylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4823	1-imidazolylmethyl	1-pyrazolylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4824	1-indolylmethyl	1-imidazolylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4825	1-triazolylmethyl	1-indolylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4826	1-tetrazolylmethyl	1-triazolylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4827	2-pyridylmethyl	1-tetrazolylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4828	3-pyridylmethyl	2-pyridylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4829	4-pyridylmethyl	3-pyridylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4830	cyclohexylmethyl	4-pyridylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4831	2-naphthylmethyl	cyclohexylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4832	3-naphthylmethyl	2-naphthylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4833	2-thiophenylmethyl	3-naphthylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl

Table 6

4834	4-(1-methyl)piperidinyl-methyl	2-thiophenylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4835	"(3,4-methylenedioxyphenyl)methyl	4-(1-methyl)piperidinyl-methyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4836	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4837	4-biphenylmethyl	2-thienylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4838	1-pyrimidinylmethyl	4-biphenylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4839	2-benzothiazolylmethyl	1-pyrimidinylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4840	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4841	2-thiomethylethyl	2-benzothiophenylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4842	2-thiomethylmethyl	2-thiomethylethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4843	2-methylpropyl	2-thiomethylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4844	2-methylbutyl	2-methylpropyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4845	3-methylbutyl	2-methylbutyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4846	cyclopropylmethyl	3-methylbutyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4847	cyclobutylmethyl	cyclopropylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4848	cyclopentylmethyl	cyclobutylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4849	p-hydroxyphenyl-methyl	cyclopentylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4850	p-nitrophenylmethyl	p-hydroxyphenyl-methyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4851	p-aminophenyl-methyl	p-nitrophenylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl

Table 6

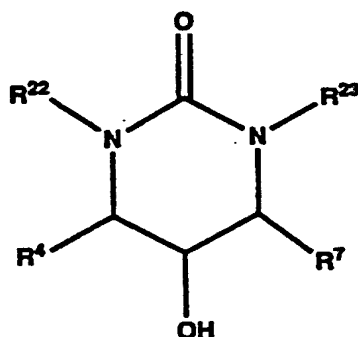
4852	"4-(N,N-dimethylamino)phenylmethyl"	P-aminophenylmethyl	2-pyridylmethyl	2-(1-imidazolyl)ethyl
4853	benzyl	"4-(N,N-dimethylamino)phenylmethyl"	2-thiomethylethyl	2-pyridylmethyl
4854	1-pyrollylmethyl	benzyl	2-thiomethylethyl	2-pyridylmethyl
4855	1-pyrazolylmethyl	1-pyrollylmethyl	2-thiomethylethyl	2-pyridylmethyl
4856	1-imidazolylmethyl	1-pyrazolylmethyl	2-thiomethylethyl	2-pyridylmethyl
4857	1-indolylmethyl	1-imidazolylmethyl	2-thiomethylethyl	2-pyridylmethyl
4858	1-triazolylmethyl	1-indolylmethyl	2-thiomethylethyl	2-pyridylmethyl
4859	1-tetrazolylmethyl	1-triazolylmethyl	2-thiomethylethyl	2-pyridylmethyl
4860	2-pyridylmethyl	1-tetrazolylmethyl	2-thiomethylethyl	2-pyridylmethyl
4861	3-pyridylmethyl	2-pyridylmethyl	2-thiomethylethyl	2-pyridylmethyl
4862	4-pyridylmethyl	3-pyridylmethyl	2-thiomethylethyl	2-pyridylmethyl
4863	cyclohexylmethyl	4-pyridylmethyl	2-thiomethylethyl	2-pyridylmethyl
4864	2-naphthylmethyl	cyclohexylmethyl	2-thiomethylethyl	2-pyridylmethyl
4865	3-naphthylmethyl	2-naphthylmethyl	2-thiomethylethyl	2-pyridylmethyl
4866	2-thiophenylmethyl	3-naphthylmethyl	2-thiomethylethyl	2-pyridylmethyl
4867	4-(1-methyl)piperidinylmethyl	1-thiophenylmethyl	2-thiomethylethyl	2-pyridylmethyl
4868	"(3,4-methylenedioxyphenyl)methyl"	4-(1-methyl)piperidinylmethyl	2-thiomethylethyl	2-pyridylmethyl

Table 6

4869	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	2-thiomethylethyl	2-pyridylmethyl
4870	4-biphenylmethyl	2-thienylmethyl	2-thiomethylethyl	2-pyridylmethyl
4871	pyrimidinylmethyl	4-biphenylmethyl	2-thiomethylethyl	2-pyridylmethyl
4872	2-benzothiazolylmethyl	pyrimidinylmethyl	2-thiomethylethyl	2-pyridylmethyl
4873	2-benzothiophenylmethyl	2-benzothiazolylmethyl	2-thiomethylethyl	2-pyridylmethyl
4874	2-thiomethylethyl	2-benzothiophenylmethyl	2-thiomethylethyl	2-pyridylmethyl
4875	2-thiomethylmethyl	2-thiomethylethyl	2-thiomethylethyl	2-pyridylmethyl
4876	2-methylpropyl	2-thiomethylmethyl	2-thiomethylethyl	2-pyridylmethyl
4877	2-methylbutyl	2-methylpropyl	2-thiomethylethyl	2-pyridylmethyl
4878	3-methylbutyl	2-methylbutyl	2-thiomethylethyl	2-pyridylmethyl
4879	cyclopropylmethyl	3-methylbutyl	2-thiomethylethyl	2-pyridylmethyl
4880	cyclobutylmethyl	cyclopropylmethyl	2-thiomethylethyl	2-pyridylmethyl
4881	cyclopentylmethyl	cyclobutylmethyl	2-thiomethylethyl	2-pyridylmethyl
4882	p-hydroxyphenylmethyl	cyclopentylmethyl	2-thiomethylethyl	2-pyridylmethyl
4883	p-nitrophenylmethyl	p-hydroxyphenylmethyl	2-thiomethylethyl	2-pyridylmethyl
4884	p-aminophenylmethyl	p-nitrophenylmethyl	2-thiomethylethyl	2-pyridylmethyl
4885	"4-(N,N-dimethylamino)phenylmethyl"	p-aminophenylmethyl	2-thiomethylethyl	2-pyridylmethyl

Table 7

TABLE 7



Example Number	R4	R7	R22	R23
4886	benzyl	benzyl	2-methallyl	2-methallyl
4887	1-pyrrolylmethyl	1-pyrrolylmethyl	4-methylpentyl	4-methylpentyl
4888	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	3-methallyl
4889	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl	3-methylbutyl
4890	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	3-methylpentyl
4891	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	3-pentyl
4892	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	4-methylpentyl
4893	4-pyridylmethyl	4-pyridylmethyl	"3,3-dimethallyl"	"3,3-dimethallyl"
4894	cyclohexylmethyl	cyclohexylmethyl	"3,3-dimethylbutyl"	"3,3-dimethylbutyl"
4895	2-thiophenylmethyl	2-thiophenylmethyl	allyl	allyl
4896	"(3,4-methylene-dioxyphenyl)" methyl	benzyl	2-methallyl	2-methallyl
4897	methyl	"(3,4-methylene-dioxyphenyl)" cyclobutylmethyl	cyclopropyl-methyl	2-methallyl
4898	2-thienylmethyl	2-thienylmethyl	cyclobutylmethyl	2-methallyl
4899			cyclopropyl-methyl	cyclopropyl-methyl
4900	4-biphenylmethyl	4-biphenylmethyl	isobutyl	isobutyl
4901	2-thiomethylethyl	2-thiomethylethyl	n-butyl	n-butyl
4902	p-hydroxyphenylmethyl	p-hydroxyphenylmethyl	n-pentyl	n-pentyl
4903	p-nitrophenylmethyl	p-nitrophenylmethyl	propyl	propyl
4904	benzyl	benzyl	2-methallyl	H
4905	1-pyrrolylmethyl	1-pyrrolylmethyl	4-methylpentyl	H

Table 7

4906	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	H
4907	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl	H
4908	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	H
4909	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	H
4910	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	H
4911	4-pyridylmethyl	4-pyridylmethyl	"3,3-dimethallyl"	H
4912	cyclohexylmethyl	cyclohexylmethyl	"3,3-dimethylbutyl"	H
4913	2-thiophenylmethyl	2-thiophenylmethyl	allyl	H
4914	"(3,4-methylenedioxyphenyl)" methyl	2-thiophenylmethyl	allyl	H
4915	methyl	"(3,4-methylene-dioxyphenyl)" methyl	allyl	H
4916	2-thienylmethyl	cyclobutylmethyl	H	H
4917		2-thienylmethyl	cyclopropyl-methyl	H
4918	4-biphenylmethyl	4-biphenylmethyl	isobutyl	H
4919	2-thiomethylethyl	2-thiomethylethyl	n-butyl	
4920	p-hydroxyphenyl-methyl	p-hydroxyphenyl-methyl	n-pentyl	H
4921	p-nitrophenylmethyl	p-nitrophenyl-methyl	propyl	H
4922	benzyl	p-nitrophenyl-methyl	2-methallyl	2-methallyl
4923	1-pyrollylmethyl	benzyl	4-methylpentyl	4-methylpentyl
4924	1-pyrazolylmethyl	1-pyrollylmethyl	3-methallyl	3-methallyl
4925	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
4926	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylpentyl
4927	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
4928	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
4929	4-pyridylmethyl	3-pyridylmethyl	"3,3-dimethallyl"	"3,3-dimethallyl"
4930	cyclohexylmethyl	4-pyridylmethyl	"3,3-dimethylbutyl"	"3,3-dimethylbutyl"
4931	2-thiophenylmethyl	cyclohexylmethyl	allyl	allyl
4932	"(3,4-methylene-dioxyphenyl)" methyl	cyclohexylmethyl	allyl	allyl
4933		2-thiophenylmethyl	cyclobutylmethyl	cyclobutyl-methyl
4934	2-thienylmethyl	"(3,4-methylene-dioxyphenyl)" methyl	cyclobutylmethyl	cyclobutyl-methyl
4935	methyl	cyclopropyl-methyl	cyclopropyl-methyl	
4936	4-biphenylmethyl	2-thienylmethyl	isobutyl	isobutyl
4937	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl

Table 7

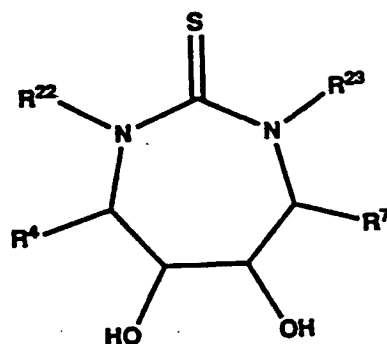
4938	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-pentyl	*
4939	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	propyl	*
4940	benzyl	p-nitrophenylmethyl	2-methallyl	2-methallyl	*
4941	1-pyrololymethyl	benzyl	4-methylpentyl	4-methylpentyl	*
4942	1-pyrazololymethyl	1-pyrololymethyl	3-methallyl	3-methallyl	
4943	1-imidazololymethyl	1-pyrazololymethyl	3-methylbutyl	3-methylbutyl	
4944	1-indololymethyl	1-imidazololymethyl	3-methylpentyl	3-methylpentyl	
4945	2-pyridolymethyl	1-indololymethyl	3-pentyl	3-pentyl	
4946	3-pyridolymethyl	2-pyridolymethyl	4-methylpentyl	4-methylpentyl	
4947	4-pyridolymethyl	3-pyridolymethyl	"3,3-dimethallyl"	"3,3-dimethallyl"	
4948	cyclohexylmethyl	4-pyridolymethyl	"3,3-dimethylbutyl"	"3,3-dimethylbutyl"	
4949	2-thiophenylmethyl	cyclohexylmethyl	allyl	allyl	
4950	"(3,4-methylene-dioxyphenyl)" methyl	4-pyridolymethyl	"3,3-dimethylbutyl"	"3,3-dimethylbutyl"	
4951	2-thienylmethyl	2-thiophenylmethyl	cyclobutylmethyl	cyclobutylmethyl	
4952	methyl	"(3,4-methylene-dioxy-phenyl)"	cyclobutylmethyl	cyclobutylmethyl	
4953	4-biphenylmethyl	cyclopropylmethyl	cyclopropylmethyl	isobutyl	
4954	2-thiophenylmethyl	2-thienylmethyl	isobutyl	isobutyl	
4955	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl	
4956	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-pentyl	
4957	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	propyl	
4958	benzyl	p-nitrophenylmethyl	2-methallyl	propyl	
4959	1-pyrololymethyl	1-benzyl	4-methylpentyl	2-methallyl	
4960	1-pyrazololymethyl	1-pyrololymethyl	3-methallyl	4-methylpentyl	
4961	1-imidazololymethyl	1-pyrazololymethyl	3-methylbutyl	3-methallyl	
4962	1-indololymethyl	1-imidazololymethyl	3-methylpentyl	3-methylbutyl	
4963	2-pyridolymethyl	1-indololymethyl	3-pentyl	3-methylpentyl	
4964	3-pyridolymethyl	2-pyridolymethyl	4-methylpentyl	3-pentyl	
4965	4-pyridolymethyl	3-pyridolymethyl	"3,3-dimethallyl"	4-methylpentyl	*
4966	cyclohexylmethyl	4-pyridolymethyl	"3,3-dimethylbutyl"	"3,3-dimethallyl"	*
4967	2-thiophenylmethyl	cyclohexylmethyl	allyl	"3,3-dimethylbutyl"	
4968	"(3,4-methylene-dioxyphenyl)"	cyclohexylmethyl	allyl	"3,3-dimethylbutyl"	*

Table 7

4969	methyl	2-thiophenylmethyl	cyclobutylmethyl	allyl
4970	2-thienylmethyl	"(3,4-methylene-dioxyphenyl)"	cyclobutylmethyl	allyl
4971	methyl	cyclopropylmethy	cyclobutylmethyl	allyl
4972	4-biphenylmethyl	1		
4973	2-thiomethylethyl	2-thienylmethyl	isobutyl	cyclopropyl-methyl
4974	p-hydroxyphenyl-methyl	4-biphenylmethyl	n-butyl	isobutyl
4975	p-nitrophenyl-methyl	2-thiomethylethyl	n-pentyl	n-butyl
4975A	benzyl	p-hydroxyphenyl-methyl	propyl	n-pentyl
		benzyl	cyclopropyl-methyl	N,N-dimethyl-aminoethyl

Table 8

TABLE 8



Example R4 Number		R7	R22	R23
4976	benzyl	benzyl	2-methallyl	2-methallyl
4977	1-pyrrolylmethyl	1-pyrrolylmethyl	4-methylpentyl	4-methylpentyl
4978	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	3-methallyl
4979	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl	3-methylbutyl
4980	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	3-methylpentyl
4981	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	3-pentyl
4982	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	4-methylpentyl
4983	4-pyridylmethyl	4-pyridylmethyl	"3,3-dimethallyl"	"3,3-dimethallyl"
4984	cyclohexylmethyl	cyclohexylmethyl	"3,3-dimethylbutyl"	"3,3-dimethylbutyl"
4985	2-thiophenylmethyl	2-thiophenylmethyl	allyl	allyl
4986	"(3,4-methylene-dioxy-phenyl)-methyl"	"(3,4-methylene-dioxyphenyl)-methyl"	cyclobutylmethyl	cyclobutylmethyl
4987	2-thienylmethyl	2-thienylmethyl	cyclopropylmethyl	cyclopropylmethyl
4988	4-biphenylmethyl	4-biphenylmethyl	isobutyl	isobutyl
4989	2-thiomethylethyl	2-thiomethylethyl	n-butyl	n-butyl
4990	p-hydroxyphenylmethyl	p-hydroxyphenylmethyl	n-pentyl	n-pentyl
4991	p-nitrophenylmethyl	p-nitrophenylmethyl	propyl	propyl

Table 8

4992	benzyl	benzyl	2-methallyl	H
4993	1-pyrolylmethyl	1-pyrolylmethyl	4-methylpentyl	H
4994	1-	1-	3-methallyl	H
	pyrazolylmethyl	pyrazolylmethyl		
4995	1-imidazolyl-	1-imidazolyl-	3-methylbutyl	H
	methyl	methyl		
4996	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	H
4997	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	H
4998	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	H
4999	4-pyridylmethyl	4-pyridylmethyl	"3,3-	H
			dimethallyl"	
5000	cyclohexyl-	cyclohexyl-	"3,3-	H
	methyl	methyl	dimethylbutyl"	
5001	2-thiophenyl-	2-thiophenyl-	allyl	H
	methyl	methyl		
5002	"(3,4-	"(3,4-	cyclobutyl-	H
	methylenedioxy-	methylenedioxy-	methyl	
	phenyl)methyl"	phenyl)methyl"		
5003	2-thienylmethyl	2-thienylmethyl	cyclopropyl-	H
			methyl	
5004	4-	4-	isobutyl	H
	biphenylmethyl	biphenylmethyl		
5005	2-	2-	n-butyl	
	thiomethylethyl	thiomethylethyl		
5006	p-	p-	n-pentyl	H
	hydroxyphenyl-	hydroxyphenyl-		
	methyl	methyl		
5007	p-nitrophenyl-	p-nitrophenyl-	propyl	H
	methyl	methyl		
5008	benzyl	p-nitrophenyl-	2-methallyl	2-methallyl
		methyl		
5009	1-pyrolylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5010	1-	1-pyrolylmethyl	3-methallyl	3-methallyl
	pyrazolylmethyl			
5011	1-imidazolyl-	1-	3-methylbutyl	3-methylbutyl
	methyl	pyrazolylmethyl		
5012	1-indolylmethyl	1-	3-methylpentyl	3-methylpentyl
		imidazolylmethy		
		l		
5013	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5014	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5015	4-pyridylmethyl	3-pyridylmethyl	"3,3-	"3,3-
			dimethallyl"	dimethallyl"
5016	cyclohexylmethy	4-pyridylmethyl	"3,3-	"3,3-
	l		dimethylbutyl"	dimethylbutyl"
5017	2-thiophenyl-	cyclohexyl-	allyl	allyl
	methyl	methyl		
5018	"(3,4-	2-thiophenyl-	cyclobutylmethy	cyclobutylmethy
	methylenedioxy-	methyl	l	l
	phenyl)methyl"			
5019	2-thienylmethyl	"(3,4-	cyclopropylmeth	cyclopropylmeth
		methylenedioxy	yl	yl
		phenyl)methyl"		
5020	4-	2-thienylmethyl	isobutyl	isobutyl
	biphenylmethyl			
5021	2-	4-	n-butyl	n-butyl
	thiomethylethyl	biphenylmethyl		

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Table 8

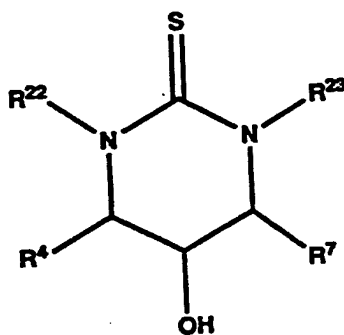
5022	p-hydroxyphenyl-methyl	2-thiomethylethyl	n-pentyl	n-pentyl	.
5023	p-nitrophenyl-methyl	p-hydroxyphenyl-methyl	propyl	propyl	.
5024	benzyl	p-nitrophenyl-methyl	2-methallyl	2-methallyl	.
5025	1-pyrrolylmethyl	benzyl	4-methylpentyl	4-methylpentyl	.
5026	1-pyrazolylmethyl	1-pyrrolylmethyl	3-methallyl	3-methallyl	.
5027	1-imidazolyl-methyl	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl	.
5028	1-indolylmethyl	1-imidazolyl-methyl	3-methylpentyl	3-methylpentyl	.
5029	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl	.
5030	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl	.
5031	4-pyridylmethyl	3-pyridylmethyl	"3,3-dimethallyl"	"3,3-dimethallyl"	.
5032	cyclohexylmethy	4-pyridylmethyl	"3,3-dimethylbutyl"	"3,3-dimethylbutyl"	.
5033	1	cyclohexylmethy	allyl	allyl	.
5034	2-thiophenyl-methyl	1	cyclobutylmethy	cyclobutylmethy	.
5035	"(3,4-methylenedioxyphenyl)methyl"	2-thiophenylmethy	1	1	.
5036	4-biphenylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclopropyl-methyl	cyclopropyl-methyl	.
5037	2-thiomethylethyl	2-thienylmethyl	isobutyl	isobutyl	.
5038	p-hydroxyphenyl-methyl	4-biphenylmethyl	n-butyl	n-butyl	.
5039	p-nitrophenyl-methyl	2-biphenylmethyl	n-pentyl	n-pentyl	.
5040	benzyl	2-thiomethylethyl	propyl	propyl	.
5041	1-pyrrolylmethyl	p-hydroxyphenylme	2-methallyl	propyl	.
5042	1-pyrazolylmethyl	thyl	4-methylpentyl	2-methallyl	.
5043	1-imidazolyl-methyl	p-nitrophenyl-methyl	3-methallyl	4-methylpentyl	.
5044	1-indolylmethyl	benzyl	3-methylbutyl	3-methallyl	.
5045	2-pyridylmethyl	1-pyrrolylmethyl	3-methylpentyl	3-methylbutyl	.
5046	3-pyridylmethyl	1-imidazolyl-methyl	3-pentyl	3-methylpentyl	.
5047	4-pyridylmethyl	pyrazolylmethyl	4-methylpentyl	3-pentyl	.
5048	cyclohexylmethy	1-indolylmethyl	"3,3-dimethallyl"	4-methylpentyl	.
5049	1	2-pyridylmethyl	"3,3-dimethylbutyl"	"3,3-dimethallyl"	.
	2-thiophenyl-methyl	3-pyridylmethyl	allyl	"3,3-dimethylbutyl"	.
		cyclohexyl-methyl			.

Table 8

5050	"(3,4-methylenedioxyphenyl)methyl"	2-thiophenylmethyl	cyclobutylmethyl	allyl
5051	2-thienylmethyl	"(3,4-methylenedioxyphenyl)methyl"	cyclopropylmethyl	cyclobutylmethyl
5052	4-biphenylmethyl	2-thienylmethyl	isobutyl	cyclopropylmethyl
5053	2-thiomethylethyl	4-biphenylmethyl	n-butyl	isobutyl
5054	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-butyl
5055	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	n-pentyl

Table 9

TABLE 9



Example R4 Number	R7	R22	R23
5056	benzyl	2-methallyl	2-methallyl
5057	1-pyrrolylmethyl	4-methylpentyl	4-methylpentyl
5058	1-pyrazolylmethyl	3-methallyl	3-methallyl
5059	1-imidazolylmethy	3-methylbutyl	3-methylbutyl
5060	1-indolylmethyl	3-methylpentyl	3-methylpentyl
5061	2-pyridylmethyl	3-pentyl	3-pentyl
5062	3-pyridylmethyl	4-methylpentyl	4-methylpentyl
5063	4-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5064	cyclohexylmethy	3,3-dimethylbutyl	3,3-dimethylbutyl
5065	1-thiophenylmethy	allyl	allyl
5066	2-(3,4-methylenedioxyphenyl)methyl	cyclobutylmethy	cyclobutylmethy
5067	2-thienylmethyl	1	1
5068	4-biphenylmethyl	cyclopropylmeth	cyclopropylmeth
5069	2-thiomethylethyl	yl	yl
5070	p-hydroxyphenylme	isobutyl	isobutyl
	thyl	n-butyl	n-butyl
		n-pentyl	n-pentyl

Table 9

			propyl	propyl
5071	p-nitrophenylmeth	p-nitrophenylmeth		
	yl	yl		
5072	benzyl	benzyl	2-methallyl	H
5073	1-pyrollylmethyl	1-pyrollylmethyl	4-methylpentyl	H
5074	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	H
5075	1-imidazolylmethy	1-imidazolylmethy	3-methylbutyl	H
	l	l		
5076	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	H
5077	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	H
5078	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	H
5079	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethallyl	H
5080	cyclohexylmethy	cyclohexylmethy	3,3-dimethylbutyl	H
	l	l	allyl	H
5081	2-thiophenylmethy	2-thiophenylmethy		
	l	l		
5082	(3,4-methylenedioxyphenyl)methyl	(3,4-methylenedioxyphenyl)methyl	cyclobutylmethy	H
	l	l		
5083	2-thienylmethyl	2-thienylmethyl	cyclopropylmeth	H
			yl	
5084	4-biphenylmethyl	4-biphenylmethyl	isobutyl	H
5085	2-thiomethylethyl	2-thiomethylethyl	n-butyl	
5086	p-hydroxyphenylme	p-hydroxyphenylme	n-pentyl	H
	thyl	thyl		
5087	p-nitrophenylmeth	p-nitrophenylmeth	propyl	H
	yl	yl		
5088	benzyl	p-nitrophenylmeth	2-methallyl	2-methallyl
		yl		
5089	1-pyrollylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5090	1-pyrazolylmethyl	1-pyrollylmethyl	3-methallyl	3-methallyl
5091	1-imidazolylmethy	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
	l			
5092	1-indolylmethyl	1-imidazolylmethy	3-methylpentyl	3-methylpentyl
		l		
5093	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5094	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5095	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5096	cyclohexylmethy	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
	l		allyl	allyl
5097	2-thiophenylmethy	cyclohexylmethy		
	l	l		

Table 9

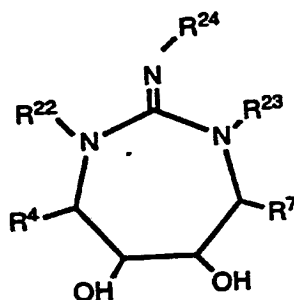
5098	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	cyclobutylmethyl
5099	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclopropylmethyl
5100	4-biphenylmethyl	2-thienylmethyl	isobutyl	isobutyl
5101	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl
5102	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-pentyl
5103	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	propyl
5104	benzyl	p-nitrophenylmethyl	2-methallyl	2-methallyl
5105	1-pyrrolylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5106	1-pyrazolylmethyl	1-pyrrolylmethyl	3-methallyl	3-methallyl
5107	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
5108	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylpentyl
5109	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5110	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5111	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5112	1-cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5113	2-thiophenylmethyl	1-cyclohexylmethyl	allyl	allyl
5114	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	cyclobutylmethyl
5115	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclopropylmethyl
5116	4-biphenylmethyl	2-thienylmethyl	isobutyl	isobutyl
5117	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl
5118	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-pentyl
5119	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	propyl
5120	benzyl	p-nitrophenylmethyl	2-methallyl	propyl
5121	1-pyrrolylmethyl	benzyl	4-methylpentyl	2-methallyl

Table 9

5122	1-pyrazolylmethyl	1-pyrollylmethyl	3-methallyl	4-methylpentyl
5123	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methallyl
5124	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylbutyl
5125	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-methylpentyl
5126	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	3-pentyl
5127	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	4-methylpentyl
5128	1-cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethallyl
5129	2-thiophenylmethyl	1-cyclohexylmethyl	allyl	3,3-dimethylbutyl
5130	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	allyl
5131	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclobutylmethyl
5132	4-biphenylmethyl	2-thienylmethyl	isobutyl	cyclopropylmethyl
5133	2-thiomethylethyl	4-biphenylmethyl	n-butyl	isobutyl
5134	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-butyl
5135	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	n-pentyl

Table 10

TABLE 10

R²⁴ = H, CH₃, or C₂H₅

<u>Example R4</u> <u>Number</u>	<u>R7</u>	<u>R22</u>	<u>R23</u>
5136	benzyl	2-methallyl	2-methallyl
5137	1-pyrrolylmethyl	4-methylpentyl	4-methylpentyl
5138	1-pyrazolylmethyl	3-methallyl	3-methallyl
5139	1-imidazolyl-methyl	3-methylbutyl	3-methylbutyl
5140	1-indolylmethyl	3-methylpentyl	3-methylpentyl
5141	2-pyridylmethyl	3-pentyl	3-pentyl
5142	3-pyridylmethyl	4-methylpentyl	4-methylpentyl
5143	4-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5144	cyclohexyl-methyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5145	2-thiophenyl-methyl	allyl	allyl
5146	(3,4-methylenedioxyphenyl)methyl	cyclobutylmethy l	cyclobutylmethy l
5147	2-thienylmethyl	cyclopropylmeth yl	cyclopropylmeth yl
5148	4-biphenyl-methyl	isobutyl	isobutyl
5149	2-thiomethylethyl	n-butyl	n-butyl
5150	p-hydroxy-phenyl-methyl	n-pentyl	n-pentyl
5151	p-nitrophenyl-methyl	propyl	propyl
5152	benzyl	2-methallyl	H
5153	1-pyrrolylmethyl	4-methylpentyl	H
5154	1-pyrazolylmethyl	3-methallyl	H

Table 10

5155	1-imidazolyl-methyl	1-imidazolyl-methyl	3-methylbutyl	H
5156	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	H
5157	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	H
5158	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	H
5159	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethylallyl	H
5160	cyclohexyl-methyl	cyclohexyl-methyl	3,3-dimethylbutyl	H
5161	2-thiophenyl-methyl	2-thiophenyl-methyl	allyl	H
5162	(3,4-methylene-dioxyphenyl)-methyl	(3,4-methylene-dioxyphenyl)-methyl	cyclobutyl-methyl	H
5163	2-thienylmethyl	2-thienylmethyl	cyclopropyl-methyl	H
5164	4-biphenylmethyl	4-biphenylmethyl	isobutyl	H
5165	2-thiomethylethyl	2-thiomethylethyl	n-butyl	
5166	p-hydroxyphenyl-methyl	p-hydroxyphenyl-methyl	n-pentyl	H
5167	p-nitrophenyl-methyl	p-nitrophenyl-methyl	propyl	H
5168	benzyl	p-nitrophenyl-methyl	2-methylallyl	2-methylallyl
5169	1-pyrollylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5170	1-pyrazolylmethyl	1-pyrollyl-methyl	3-methylallyl	3-methylallyl
5171	1-imidazolylmethy	1-pyrazolyl-methyl	3-methylbutyl	3-methylbutyl
5172	1-indolylmethyl	1-imidazolyl-methyl	3-methylpentyl	3-methylpentyl
5173	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5174	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5175	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylallyl	3,3-dimethylallyl
5176	cyclohexyl-methyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5177	2-thiophenyl-methyl	cyclohexyl-methyl	allyl	allyl
5178	(3,4-methylenedioxyphenyl)methyl	2-thiophenyl-methyl	cyclobutylmethy	cyclobutylmethy
5179	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	1	1
5180	4-biphenylmethyl	2-thienylmethyl	cyclopropylmeth	cyclopropylmeth
5181	2-thiomethylethyl	4-biphenylmethyl	yl	yl
5182	p-hydroxyphenyl-methyl	2-thiomethylethyl	isobutyl	isobutyl
			n-butyl	n-butyl
			n-pentyl	n-pentyl

Table 10

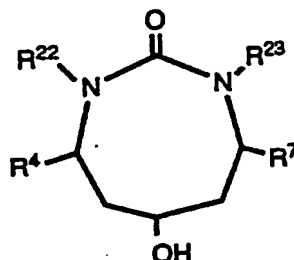
5183	p-nitrophenyl-methyl	p-hydroxyphenyl-methyl	propyl	propyl
5184	benzyl	p-nitrophenyl-methyl	2-methallyl	2-methallyl
5185	1-pyrolylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5186	1-pyrazolylmethyl	1-pyrolylmethyl	3-methallyl	3-methallyl
5187	1-imidazolyl-methyl	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
5188	1-indolylmethyl	1-imidazolylmethy l	3-methylpentyl	3-methylpentyl
5189	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5190	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5191	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5192	cyclohexyl-methyl	4-pyridylmethyl	3,3-dimethylbutyl allyl-	3,3-dimethylbutyl allyl
5193	2-thiophenyl-methyl	cyclohexyl-methyl	cyclobutylmethy l	cyclobutyl-methyl
5194	(3,4-methylenedioxy-phenyl)methyl	2-thiophenyl-methyl	cyclopropyl-methyl	cyclopropyl-methyl
5195	2-thienylmethyl	(3,4-methylenedioxy-phenyl)methyl	isobutyl	isobutyl
5196	4-biphenylmethyl	2-thienylmethyl	n-butyl	n-butyl
5197	2-thiomethylethyl	4-biphenylmethyl	n-pentyl	n-pentyl
5198	p-hydroxyphenyl-methyl	2-thiomethylethyl	propyl	propyl
5199	p-nitrophenyl-methyl	p-hydroxyphenyl-methyl	2-methallyl	propyl
5200	benzyl	p-nitrophenyl-methyl	4-methylpentyl	2-methallyl
5201	1-pyrolylmethyl	benzyl	3-methallyl	4-methylpentyl
5202	1-pyrazolylmethyl	1-pyrolylmethyl	3-methylbutyl	3-methallyl
5203	1-imidazolyl-methyl	1-pyrazolylmethyl	3-methylpentyl	3-methylbutyl
5204	1-indolylmethyl	1-imidazolyl-methyl	3-pentyl	3-methylpentyl
5205	2-pyridylmethyl	1-indolylmethyl	4-methylpentyl	3-pentyl
5206	3-pyridylmethyl	2-pyridylmethyl	3,3-dimethallyl	4-methylpentyl
5207	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylbutyl allyl	3,3-dimethallyl
5208	cyclohexyl-methyl	4-pyridylmethyl	cyclobutylmethy l	3,3-dimethylbutyl allyl
5209	2-thiophenyl-methyl	cyclohexyl-methyl		
5210	(3,4-methylenedioxyphenyl)methyl	2-thiophenyl-methyl		

Table 10

5211	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclobutylmethyl
5212	4-biphenylmethyl	2-thienylmethyl	isobutyl	cyclopropylmethyl
5213	2-thiomethylethyl	4-biphenylmethyl	n-butyl	isobutyl
5214	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-butyl
5215	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	n-pentyl

Table 11

Table 11



<u>Example R4</u> <u>Number</u>		<u>R7</u>	<u>R22</u>	<u>R23</u>
5216	benzyl	benzyl	2-methallyl	2-methallyl
5217	1-pyrrolylmethyl	1-pyrrolylmethyl	4-methylpentyl	4-methylpentyl
5218	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	3-methallyl
5219	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl	3-methylbutyl
5220	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	3-methylpentyl
5221	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	3-pentyl
5222	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	4-methylpentyl
5223	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5224	cyclohexylmethyl	cyclohexylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5225	2-thiophenylmethyl	2-thiophenylmethyl	allyl	allyl
5226	(3,4-methylene-dioxyphenyl)-methyl	(3,4-methylene-dioxyphenyl)-methyl	cyclobutylmethyl	cyclobutylmethyl
5227	2-thienylmethyl	2-thienylmethyl	cyclopropylmethyl	cyclopropylmethyl
5228	4-biphenylmethyl	4-biphenylmethyl	isobutyl	isobutyl
5229	2-thiomethylethyl	2-thiomethylethyl	n-butyl	n-butyl
5230	p-hydroxyphenylmethyl	p-hydroxyphenylmethyl	n-pentyl	n-pentyl
5231	p-nitrophenylmethyl	p-nitrophenylmethyl	propyl	propyl
5232	benzyl	benzyl	2-methallyl	H
5233	1-pyrrolylmethyl	1-pyrrolylmethyl	4-methylpentyl	H
5234	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	H
5235	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl	H

Table 11

5236	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	H
5237	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	H
5238	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	H
5239	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethylallyl	H
5240	cyclohexyl-methyl	cyclohexyl-methyl	3,3-dimethylbutyl	H
5241	2-thiophenyl-methyl	2-thiophenyl-methyl	allyl	H
5242	(3,4-methylene-dioxyphenyl)-methyl	(3,4-methylene-dioxyphenyl)-methyl	cyclobutyl-methyl	H
5243	2-thienylmethyl	2-thienylmethyl	cyclopropyl-methyl	H
5244	4-biphenylmethyl	4-biphenylmethyl	isobutyl	H
5245	2-thiomethylethyl	2-thiomethylethyl	n-butyl	
5246	p-hydroxyphenyl-methyl	p-hydroxyphenyl-methyl	n-pentyl	H
5247	p-nitrophenyl-methyl	p-nitrophenyl-methyl	propyl	H
5248	benzyl	p-nitrophenyl-methyl	2-methylallyl	2-methylallyl
5249	1-pyrollylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5250	1-pyrazolylmethyl	1-pyrollylmethyl	3-methylallyl	3-methylallyl
5251	1-imidazolyl-methyl	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
5252	1-indolylmethyl	1-imidazolyl-methyl	3-methylpentyl	3-methylpentyl
5253	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5254	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5255	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylallyl	3,3-dimethylallyl
5256	cyclohexyl-methyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5257	2-thiophenyl-methyl	cyclohexyl-methyl	allyl	allyl
5258	(3,4-methylenedioxyphenyl)methyl	2-thiophenyl-methyl	cyclobutyl-methyl	cyclobutyl-methyl
5259	2-thienylmethyl	(3,4-methylene-dioxyphenyl)-methyl	cyclopropyl-methyl	cyclopropyl-methyl
5260	4-biphenylmethyl	2-thienylmethyl	isobutyl	isobutyl
5261	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl
5262	p-hydroxyphenyl-methyl	2-thiomethylethyl	n-pentyl	n-pentyl
5263	p-nitrophenyl-methyl	p-hydroxyphenyl-methyl	propyl	propyl
5264	benzyl	p-nitrophenyl-methyl	2-methylallyl	2-methylallyl

Table 11

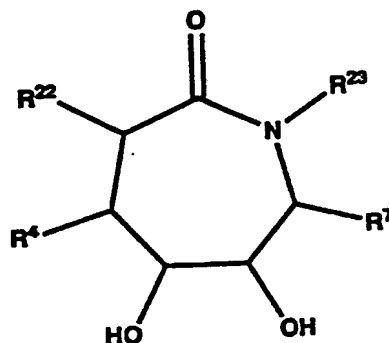
5265	1-pyrolylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5266	1-pyrazolylmethyl	1-pyrolylmethyl	3-methallyl	3-methallyl
5267	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
5268	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylpentyl
5269	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5270	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5271	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5272	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5273	2-thiophenylmethyl	cyclohexylmethyl	allyl	allyl
5274	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	cyclobutylmethyl
5275	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclopropylmethyl
5276	4-biphenylmethyl	2-thienylmethyl	isobutyl	isobutyl
5277	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl
5278	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-pentyl
5279	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	propyl
5280	benzyl	p-nitrophenylmethyl	2-methallyl	propyl
5281	1-pyrolylmethyl	benzyl	4-methylpentyl	2-methallyl
5282	1-pyrazolylmethyl	1-pyrolylmethyl	3-methallyl	4-methylpentyl
5283	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methallyl
5284	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylbutyl
5285	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-methylpentyl
5286	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	3-pentyl
5287	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	4-methylpentyl
5288	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethallyl
5289	2-thiophenylmethyl	cyclohexylmethyl	allyl	3,3-dimethylbutyl
5290	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	allyl
5291	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclobutylmethyl
5292	4-biphenylmethyl	2-thienylmethyl	isobutyl	cyclopropylmethyl

Table 11

5293	2- thiomethylethyl	4- biphenylmethyl	n-butyl	isobutyl
5294	p- hydroxyphenyl- methyl	2- thiomethylethyl	n-pentyl	n-butyl
5295	p-nitrophenyl- methyl	p-- hydroxyphenylme thyl	propyl	n-pentyl

Table 12

TABLE 12



<u>Example R4</u> <u>Number</u>		<u>R7</u>	<u>R22</u>	<u>R23</u>
5296	benzyl	benzyl	2-methallyl	2-methallyl
5297	1-pyrolylmethyl	1-pyrolylmethyl	4-methylpentyl	4-methylpentyl
5298	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	3-methallyl
5299	1-imidazolylmethy	1-imidazolylmethy	3-methylbutyl	3-methylbutyl
5300	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	3-methylpentyl
5301	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	3-pentyl
5302	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	4-methylpentyl
5303	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5304	cyclohexylmethy	cyclohexylmethy	3,3-dimethylbutyl	3,3-dimethylbutyl
5305	1-thiophenylmethy	1-thiophenylmethy	allyl	allyl
5306	(3,4-methylenedioxyphenyl)methyl	(3,4-methylenedioxyphenyl)methyl	cyclobutylmethy	cyclobutylmethy
5307	2-thienylmethyl	2-thienylmethyl	1-cyclopropylmeth	1-cyclopropylmeth
5308	4-biphenylmethyl	4-biphenylmethyl	yl isobutyl	yl isobutyl
5309	2-thiomethylethyl	2-thiomethylethyl	n-butyl	n-butyl
5310	p-hydroxyphenylme	p-hydroxyphenylme	n-pentyl	n-pentyl
	thyl	thyl		

Table 12

5311	p-nitrophenylmeth yl	p-nitrophenylmeth yl	propyl	propyl
5312	benzyl	benzyl	2-methallyl	H
5313	1-pyrollylmethyl	1-pyrollylmethyl	4-methylpentyl	H
5314	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	H
5315	1-imidazolylmethy l	1-imidazolylmethy l	3-methylbutyl	H
5316	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	H
5317	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	H
5318	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	H
5319	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethallyl	H
5320	cyclohexylmethy l	cyclohexylmethy l	3,3-dimethylbutyl	H
5321	2-thiophenylmethy l	2-thiophenylmethy l	allyl	H
5322	(3,4-methylenedioxy phenyl)methyl	(3,4-methylenedioxy phenyl)methyl	cyclobutylmethy l	H
5323	2-thienylmethyl	2-thienylmethyl	cyclopropylmeth yl	H
5324	4-biphenylmethyl	4-biphenylmethyl	isobutyl	H
5325	2-thiomethylethyl	2-thiomethylethyl	n-butyl	
5326	p-hydroxyphenylme thyl	p-hydroxyphenylme thyl	n-pentyl	H
5327	p-nitrophenylmeth yl	p-nitrophenylmeth yl	propyl	H
5328	benzyl	p-nitrophenylmeth yl	2-methallyl	2-methallyl
5329	1-pyrollylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5330	1-pyrazolylmethyl	1-pyrollylmethyl	3-methallyl	3-methallyl
5331	1-imidazolylmethy l	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
5332	1-indolylmethyl	1-imidazolylmethy l	3-methylpentyl	3-methylpentyl
5333	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5334	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5335	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5336	cyclohexylmethy l	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5337	2-thiophenylmethy l	cyclohexylmethy l	allyl	allyl

Table 12

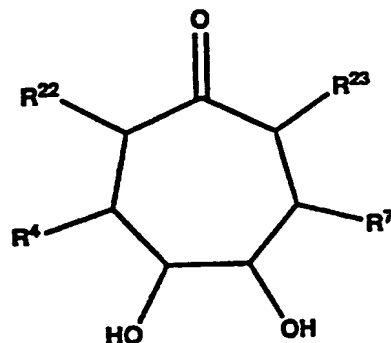
5338	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	cyclobutylmethyl	e
5339	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclopropylmethyl	a
5340	4-biphenylmethyl	2-thienylmethyl	isobutyl	isobutyl	"
5341	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl	"
5342	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-pentyl	"
5343	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	propyl	"
5344	benzyl	p-nitrophenylmethyl	2-methallyl	2-methallyl	"
5345	1-pyrolylmethyl	benzyl	4-methylpentyl	4-methylpentyl	"
5346	1-pyrazolylmethyl	1-pyrolylmethyl	3-methallyl	3-methallyl	"
5347	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl	"
5348	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylpentyl	"
5349	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl	"
5350	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl	"
5351	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl	"
5352	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl	"
5353	1-cyclohexylmethyl	1-cyclohexylmethyl	allyl	allyl	"
5354	2-thiophenylmethyl	2-thiophenylmethyl	cyclobutylmethyl	cyclobutylmethyl	"
5355	(3,4-methylenedioxyphenyl)methyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclopropylmethyl	"
5356	2-thienylmethyl	2-thienylmethyl	isobutyl	isobutyl	"
5357	4-biphenylmethyl	4-biphenylmethyl	n-butyl	n-butyl	"
5358	2-thiomethylethyl	2-thiomethylethyl	n-pentyl	n-pentyl	"
5359	p-hydroxyphenylmethyl	p-hydroxyphenylmethyl	propyl	propyl	"
5360	p-nitrophenylmethyl	p-nitrophenylmethyl	2-methallyl	propyl	"
5361	benzyl	benzyl	4-methylpentyl	2-methallyl	"

Table 12

5362	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	4-methylpentyl
5363	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methallyl
5364	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylbutyl
5365	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-methylpentyl
5366	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	3-pentyl
5367	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	4-methylpentyl
5368	1-cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethallyl
5369	2-thiophenylmethyl	1-cyclohexylmethyl	allyl	3,3-dimethylbutyl
5370	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	allyl
5371	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclobutylmethyl
5372	4-biphenylmethyl	2-thienylmethyl	isobutyl	cyclopropylmethyl
5373	2-thiomethylethyl	4-biphenylmethyl	n-butyl	isobutyl
5374	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-butyl
5375	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	n-pentyl

Table 13

TABLE 13



<u>Example</u>	<u>R4</u>	<u>R7</u>	<u>R22</u>	<u>R23</u>
5376	benzyl	benzyl	2-methallyl	2-methallyl
5377	1-pyrolylmethyl	1-pyrolylmethyl	4-methylpentyl	4-methylpentyl
5378	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	3-methallyl
5379	1-imidazolylmethy	1-imidazolylmethy	3-methylbutyl	3-methylbutyl
5380	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	3-methylpentyl
5381	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	3-pentyl
5382	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	4-methylpentyl
5383	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5384	cyclohexylmethy	cyclohexylmethy	3,3-dimethylbutyl	3,3-dimethylbutyl
5385	1-2-thiophenylmethy	1-2-thiophenylmethy	allyl	allyl
5386	(3,4-methylenedioxyphenyl)methyl	(3,4-methylenedioxyphenyl)methyl	cyclobutylmethy	cyclobutylmethy
5387	2-thienylmethyl	2-thienylmethyl	1-cyclopropylmeth	1-cyclopropylmeth
5388	4-biphenylmethyl	4-biphenylmethyl	yl isobutyl	yl isobutyl
5389	2-thiomethylethyl	2-thiomethylethyl	n-butyl	n-butyl
5390	p-hydroxyphenylme	p-hydroxyphenylme	n-pentyl	n-pentyl
	thyl	thyl		

Table 13

			propyl	propyl
5391	p-nitrophenylmeth yl	p-nitrophenylmeth yl		
5392	benzyl	benzyl	2-methallyl	H
5393	1-pyrolylmethyl	1-pyrolylmethyl	4-methylpentyl	H
5394	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl	H
5395	1-imidazolylmethy l	1-imidazolylmethy l	3-methylbutyl	H
5396	1-indolylmethyl	1-indolylmethyl	3-methylpentyl	H
5397	2-pyridylmethyl	2-pyridylmethyl	3-pentyl	H
5398	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl	H
5399	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethallyl	H
5400	cyclohexylmethy l	cyclohexylmethy l	3,3-dimethylbutyl	H
5401	2-thiophenylmethy l	2-thiophenylmethy l	allyl	H
5402	(3,4-methylenedioxy phenyl)methyl	(3,4-methylenedioxy phenyl)methyl	cyclobutylmethy l	H
5403	2-thienylmethyl	2-thienylmethyl	cyclopropylmeth yl	H
5404	4-biphenylmethyl	4-biphenylmethyl	isobutyl	H
5405	2-thiomethylethyl	2-thiomethylethyl	n-butyl	
5406	p-hydroxyphenylme thyl	p-hydroxyphenylme thyl	n-pentyl	H
5407	p-nitrophenylmeth yl	p-nitrophenylmeth yl	propyl	H
5408	benzyl	p-nitrophenylmeth yl	2-methallyl	2-methallyl
5409	1-pyrolylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5410	1-pyrazolylmethyl	1-pyrolylmethyl	3-methallyl	3-methallyl
5411	1-imidazolylmethy l	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
5412	1-indolylmethyl	1-imidazolylmethy l	3-methylpentyl	3-methylpentyl
5413	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5414	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5415	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5416	cyclohexylmethy l	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5417	2-thiophenylmethy l	cyclohexylmethy l	allyl	allyl

Table 13

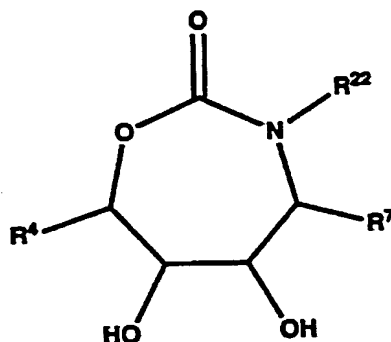
5418	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	cyclobutylmethyl
5419	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclopropylmethyl
5420	4-biphenylmethyl	2-thienylmethyl	isobutyl	isobutyl
5421	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl
5422	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-pentyl
5423	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	propyl
5424	benzyl	p-nitrophenylmethyl	2-methallyl	2-methallyl
5425	1-pyrrolylmethyl	benzyl	4-methylpentyl	4-methylpentyl
5426	1-pyrazolylmethyl	1-pyrrolylmethyl	3-methallyl	3-methallyl
5427	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methylbutyl
5428	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylpentyl
5429	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-pentyl
5430	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	4-methylpentyl
5431	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	3,3-dimethallyl
5432	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethylbutyl
5433	1-cyclohexylmethyl	1-cyclohexylmethyl	allyl	allyl
5434	2-thiophenylmethyl	1-cyclohexylmethyl	cyclobutylmethyl	cyclobutylmethyl
5435	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	1-cyclobutylmethyl	1-cyclobutylmethyl
5436	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclopropylmethyl
5437	4-biphenylmethyl	2-thienylmethyl	isobutyl	isobutyl
5438	2-thiomethylethyl	4-biphenylmethyl	n-butyl	n-butyl
5439	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-pentyl
5440	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	propyl
5441	benzyl	p-nitrophenylmethyl	2-methallyl	propyl
	1-pyrrolylmethyl	benzyl	4-methylpentyl	2-methallyl

Table 13

5442	1-pyrazolylmethyl	1-pyrollylmethyl	3-methallyl	4-methylpentyl
5443	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl	3-methallyl
5444	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl	3-methylbutyl
5445	2-pyridylmethyl	1-indolylmethyl	3-pentyl	3-methylpentyl
5446	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl	3-pentyl
5447	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethallyl	4-methylpentyl
5448	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl	3,3-dimethallyl
5449	2-thiophenylmethyl	cyclohexylmethyl	allyl	3,3-dimethylbutyl
5450	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl	allyl
5451	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl	cyclobutylmethyl
5452	4-biphenylmethyl	2-thienylmethyl	isobutyl	cyclopropylmethyl
5453	2-thiomethylethyl	4-biphenylmethyl	n-butyl	isobutyl
5454	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl	n-butyl
5455	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl	n-pentyl

Table 14

TABLE 14



Example R4
Number

R7

R22

5456	benzyl	benzyl	2-methallyl
5457	1-pyrolylmethyl	1-pyrolylmethyl	4-methylpentyl
5458	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl
5459	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl
5460	1-indolylmethyl	1-indolylmethyl	3-methylpentyl
5461	2-pyridylmethyl	2-pyridylmethyl	3-pentyl
5462	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl
5463	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethallyl
5464	cyclohexylmethyl	cyclohexylmethyl	3,3-dimethylbutyl
5465	2-thiophenylmethyl	2-thiophenylmethyl	allyl
5466	(3,4-methylenedioxyphenyl)methyl	(3,4-methylenedioxyphenyl)methyl	cyclobutylmethyl
5467	2-thienylmethyl	2-thienylmethyl	cyclopropylmethyl
5468	4-biphenylmethyl	4-biphenylmethyl	isobutyl
5469	2-thiomethylethyl	2-thiomethylethyl	n-butyl
5470	p-hydroxyphenylmethyl	p-hydroxyphenylmethyl	n-pentyl
5471	p-nitrophenylmethyl	p-nitrophenylmethyl	propyl
5472	benzyl	benzyl	2-methallyl
5473	1-pyrolylmethyl	1-pyrolylmethyl	4-methylpentyl
5474	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl
5475	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl

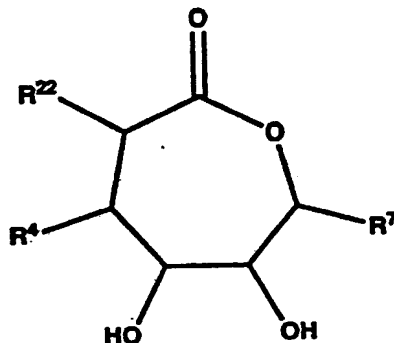
Table 14

5476	1-indolylmethyl	1-indolylmethyl	3-methylpentyl
5477	2-pyridylmethyl	2-pyridylmethyl	3-pentyl
5478	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl
5479	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethylallyl
5480	cyclohexylmethyl	cyclohexylmethyl	3,3-dimethylbutyl
5481	2-thiophenylmethyl	2-thiophenylmethyl	allyl
5482	(3,4-methylenedioxyphenyl)methyl	(3,4-methylenedioxyphenyl)methyl	cyclobutylmethyl
5483	2-thienylmethyl	2-thienylmethyl	cyclopropylmethyl
5484	4-biphenylmethyl	4-biphenylmethyl	isobutyl
5485	2-thiomethylethyl	2-thiomethylethyl	n-butyl
5486	p-hydroxyphenylmethyl	p-hydroxyphenylmethyl	n-pentyl
5487	p-nitrophenylmethyl	p-nitrophenylmethyl	propyl
5488	benzyl	p-nitrophenylmethyl	2-methylallyl
5489	1-pyrrolylmethyl	benzyl	4-methylpentyl
5490	1-pyrazolylmethyl	1-pyrrolylmethyl	3-methylallyl
5491	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl
5492	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl
5493	2-pyridylmethyl	1-indolylmethyl	3-pentyl
5494	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl
5495	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylallyl
5496	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl
5497	2-thiophenylmethyl	cyclohexylmethyl	allyl
5498	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl
5499	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl
5500	4-biphenylmethyl	2-thienylmethyl	isobutyl
5501	2-thiomethylethyl	4-biphenylmethyl	n-butyl
5502	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl
5503	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl
5504	benzyl	p-nitrophenylmethyl	2-methylallyl
5505	1-pyrrolylmethyl	benzyl	4-methylpentyl
5506	1-pyrazolylmethyl	1-pyrrolylmethyl	3-methylallyl
5507	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl
5508	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl

		Table 14	
5509	2-pyridylmethyl	1-indolylmethyl	3-pentyl
5510	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl
5511	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylallyl
5512	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl
5513	2-thiophenylmethyl	cyclohexylmethyl	allyl
5514	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl
5515	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl
5516	4-biphenylmethyl	2-thienylmethyl	isobutyl
5517	2-thiomethylethyl	4-biphenylmethyl	n-butyl
5518	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl
5519	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl
5520	benzyl	p-nitrophenylmethyl	2-methylallyl
5521	1-pyrollylmethyl	benzyl	4-methylpentyl
5522	1-pyrazolylmethyl	1-pyrollylmethyl	3-methylallyl
5523	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl
5524	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl
5525	2-pyridylmethyl	1-indolylmethyl	3-pentyl
5526	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl
5527	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylallyl
5528	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl
5529	2-thiophenylmethyl	cyclohexylmethyl	allyl
5530	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl
5531	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl
5532	4-biphenylmethyl	2-thienylmethyl	isobutyl
5533	2-thiomethylethyl	4-biphenylmethyl	n-butyl
5534	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl
5535	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl

Table 15

TABLE 15



Example R4
Number

R7

R22

5536	benzyl	benzyl	2-methallyl
5537	1-pyrrolylmethyl	1-pyrrolylmethyl	4-methylpentyl
5538	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl
5539	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl
5540	1-indolylmethyl	1-indolylmethyl	3-methylpentyl
5541	2-pyridylmethyl	2-pyridylmethyl	3-pentyl
5542	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl
5543	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethallyl
5544	cyclohexylmethyl	cyclohexylmethyl	3,3-dimethylbutyl
5545	2-thiophenylmethyl	2-thiophenylmethyl	allyl
5546	(3,4-methylenedioxyphenyl)methyl	(3,4-methylenedioxyphenyl)methyl	cyclobutylmethyl
5547	2-thienylmethyl	2-thienylmethyl	cyclopropylmethyl
5548	4-biphenylmethyl	4-biphenylmethyl	isobutyl
5549	2-thiomethylethyl	2-thiomethylethyl	n-butyl
5550	p-hydroxyphenylmethyl	p-hydroxyphenylmethyl	n-pentyl
5551	p-nitrophenylmethyl	p-nitrophenylmethyl	propyl
5552	benzyl	benzyl	2-methallyl
5553	1-pyrrolylmethyl	1-pyrrolylmethyl	4-methylpentyl
5554	1-pyrazolylmethyl	1-pyrazolylmethyl	3-methallyl
5555	1-imidazolylmethyl	1-imidazolylmethyl	3-methylbutyl

Table 15

5556	1-indolylmethyl	1-indolylmethyl	3-methylpentyl
5557	2-pyridylmethyl	2-pyridylmethyl	3-pentyl
5558	3-pyridylmethyl	3-pyridylmethyl	4-methylpentyl
5559	4-pyridylmethyl	4-pyridylmethyl	3,3-dimethylallyl
5560	cyclohexylmethyl	cyclohexylmethyl	3,3-dimethylbutyl
5561	2-thiophenylmethyl	2-thiophenylmethyl	allyl
5562	(3,4-methylenedioxyphenyl)methyl	(3,4-methylenedioxyphenyl)methyl	cyclobutylmethyl
5563	2-thienylmethyl	2-thienylmethyl	cyclopropylmethyl
5564	4-biphenylmethyl	4-biphenylmethyl	isobutyl
5565	2-thiomethylethyl	2-thiomethylethyl	n-butyl
5566	p-hydroxyphenylmethyl	p-hydroxyphenylmethyl	n-pentyl
5567	p-nitrophenylmethyl	p-nitrophenylmethyl	propyl
5568	benzyl	p-nitrophenylmethyl	2-methylallyl
5569	1-pyrollylmethyl	benzyl	4-methylpentyl
5570	1-pyrazolylmethyl	1-pyrollylmethyl	3-methylallyl
5571	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl
5572	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl
5573	2-pyridylmethyl	1-indolylmethyl	3-pentyl
5574	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl
5575	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylallyl
5576	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl
5577	2-thiophenylmethyl	cyclohexylmethyl	allyl
5578	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl
5579	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl
5580	4-biphenylmethyl	2-thienylmethyl	isobutyl
5581	2-thiomethylethyl	4-biphenylmethyl	n-butyl
5582	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl
5583	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl
5584	benzyl	p-nitrophenylmethyl	2-methylallyl
5585	1-pyrollylmethyl	benzyl	4-methylpentyl
5586	1-pyrazolylmethyl	1-pyrollylmethyl	3-methylallyl
5587	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl
5588	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl

Table 15

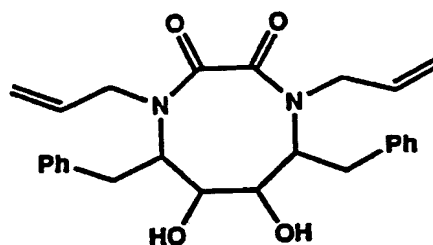
5589	2-pyridylmethyl	1-indolylmethyl	3-pentyl
5590	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl
5591	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylallyl
5592	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl
5593	2-thiophenylmethyl	cyclohexylmethyl	allyl
5594	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl
5595	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl
5596	4-biphenylmethyl	2-thienylmethyl	isobutyl
5597	2-thiomethylethyl	4-biphenylmethyl	n-butyl
5598	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl
5599	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl
5600	benzyl	p-nitrophenylmethyl	2-methylallyl
5601	1-pyrrolylmethyl	benzyl	4-methylpentyl
5602	1-pyrazolylmethyl	1-pyrrolylmethyl	3-methylallyl
5603	1-imidazolylmethyl	1-pyrazolylmethyl	3-methylbutyl
5604	1-indolylmethyl	1-imidazolylmethyl	3-methylpentyl
5605	2-pyridylmethyl	1-indolylmethyl	3-pentyl
5606	3-pyridylmethyl	2-pyridylmethyl	4-methylpentyl
5607	4-pyridylmethyl	3-pyridylmethyl	3,3-dimethylallyl
5608	cyclohexylmethyl	4-pyridylmethyl	3,3-dimethylbutyl
5609	2-thiophenylmethyl	cyclohexylmethyl	allyl
5610	(3,4-methylenedioxyphenyl)methyl	2-thiophenylmethyl	cyclobutylmethyl
5611	2-thienylmethyl	(3,4-methylenedioxyphenyl)methyl	cyclopropylmethyl
5612	4-biphenylmethyl	2-thienylmethyl	isobutyl
5613	2-thiomethylethyl	4-biphenylmethyl	n-butyl
5614	p-hydroxyphenylmethyl	2-thiomethylethyl	n-pentyl
5615	p-nitrophenylmethyl	p-hydroxyphenylmethyl	propyl

Table 16

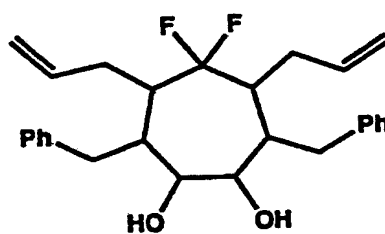
TABLE 16

**Example
Number**

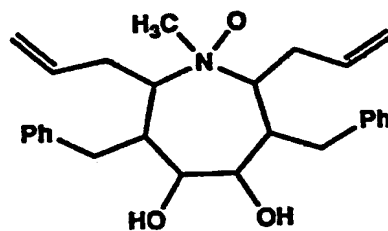
5616



5617



5618



5619

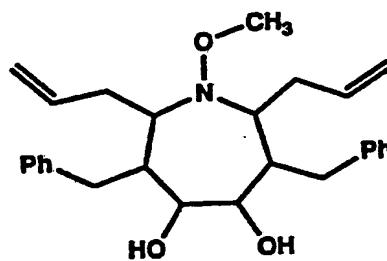
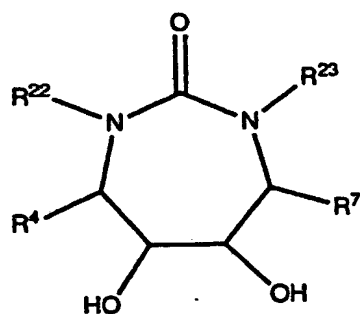


Table 17



Ex. No.	R ²²	R ²³	R ⁴ = R ⁷
5620	(m-hydroxymethyl)benzyl	(m-tetrazolyl)benzyl	benzyl
5621	(p-hydroxymethyl)benzyl	(m-tetrazolyl)benzyl	benzyl
5622	(m-hydroxy)benzyl	(m-tetrazolyl)benzyl	benzyl
5623	(p-hydroxy)benzyl	(m-tetrazolyl)benzyl	benzyl
5624	(m-carboxamido)benzyl	(m-tetrazolyl)benzyl	benzyl
5625	(m-(N-methyl carboxamido))benzyl	(m-tetrazolyl)benzyl	benzyl
5626	(m-hydroxymethyl)benzyl	(m-imidazolyl)benzyl	benzyl
5627	(p-hydroxymethyl)benzyl	(m-imidazolyl)benzyl	benzyl
5628	(m-hydroxy)benzyl	(m-imidazolyl)benzyl	benzyl
5629	(p-hydroxy)benzyl	(m-imidazolyl)benzyl	benzyl
5630	(m-carboxamido)benzyl	(m-imidazolyl)benzyl	benzyl
5631	(m-(N-methyl carboxamido))benzyl	(m-imidazolyl)benzyl	benzyl
5632	(m-hydroxymethyl)benzyl	(m-tetrazolyl)benzyl	(p-fluoro)benzyl
5633	(p-hydroxymethyl)benzyl	(m-tetrazolyl)benzyl	(p-fluoro)benzyl
5634	(m-hydroxy)benzyl	(m-tetrazolyl)benzyl	(p-fluoro)benzyl
5635	(p-hydroxy)benzyl	(m-tetrazolyl)benzyl	(p-fluoro)benzyl
5636	(m-carboxamido)benzyl	(m-tetrazolyl)benzyl	(p-fluoro)benzyl
5637	(m-(N-methyl carboxamido))benzyl	(m-tetrazolyl)benzyl	(p-fluoro)benzyl
5638	(m-hydroxymethyl)benzyl	(m-imidazolyl)benzyl	(p-fluoro)benzyl
5639	(p-hydroxymethyl)benzyl	(m-imidazolyl)benzyl	(p-fluoro)benzyl
5640	(m-hydroxy)benzyl	(m-imidazolyl)benzyl	(p-fluoro)benzyl

5641	(p-hydroxy)benzyl	(m-imidazolyl)benzyl	(p-fluoro)benzyl
5642	(m-carboxamido)benzyl	(m-imidazolyl)benzyl	(p-fluoro)benzyl
5643	(m-(N-methyl carboxamido))benzyl	(m-imidazolyl)benzyl	(p-fluoro)benzyl
5644	(m-carboxy)benzyl	(m-hydroxy)benzyl	benzyl
5645	(m-carboxy)benzyl	(p-hydroxy)benzyl	benzyl
5646	(m-carboxy)benzyl	(m-hydroxymethyl) benzyl	benzyl
5647	(m-carboxy)benzyl	(p-hydroxymethyl) benzyl	benzyl
5648	(m-carboxy)benzyl	(m-carboxamido) benzyl	benzyl
5649	(m-carboxy)benzyl	(m-(N-methyl carboxamido))benzyl	benzyl
5650	(m-carboxy)benzyl	(m-acetyl)benzyl	benzyl
5651	(m-carboxy)benzyl	(m-glycolyl)benzyl	benzyl
5652	(m-carboxy)benzyl	(m-sulfonamido) benzyl	benzyl
5653	(m-carboxy)benzyl	(m-(N-methylamino) benzyl	benzyl
5654	(m-carboxamido)benzyl	(m-hydroxy)benzyl	benzyl
5655	(m-carboxamido)benzyl	(p-hydroxy)benzyl	benzyl
5656	(m-carboxamido)benzyl	(m-hydroxymethyl) benzyl	benzyl
5657	(m-carboxamido)benzyl	(p-hydroxymethyl) benzyl	benzyl
5658	(m-carboxamido)benzyl	(m-carboxamido) benzyl	benzyl
5659	(m-carboxamido)benzyl	(m-(N-methyl carboxamido))benzyl	benzyl
5660	(m-carboxamido)benzyl	(m-acetyl)benzyl	benzyl
5661	(m-carboxamido)benzyl	(m-glycolyl)benzyl	benzyl
5662	(m-carboxamido)benzyl	(m-sulfonamido) benzyl	benzyl
5663	(m-carboxamido)benzyl	(m-(N-methylamino) benzyl	benzyl

5664	(m- (N-methyl carboxamido)) benzyl	(m-hydroxy) benzyl	benzyl
5665	(m- (N-methyl carboxamido)) benzyl	(p-hydroxy) benzyl	benzyl
5666	(m- (N-methyl carboxamido)) benzyl	(m-hydroxymethyl) benzyl	benzyl
5667	(m- (N-methyl carboxamido)) benzyl	(p-hydroxymethyl) benzyl	benzyl
5668	(m- (N-methyl carboxamido)) benzyl	(m-carboxamido) benzyl	benzyl
5669	(m- (N-methyl carboxamido)) benzyl	(m- (N-methyl carboxamido)) benzyl	benzyl
5670	(m- (N-methyl carboxamido)) benzyl	(m-acetyl) benzyl	benzyl
5671	(m- (N-methyl carboxamido)) benzyl	(m-glycolyl) benzyl	benzyl
5672	(m- (N-methyl carboxamido)) benzyl	(m-sulfonamido) benzyl	benzyl
5673	(m- (N-methyl carboxamido)) benzyl	(m- (N-methylamino) benzyl	benzyl
5674	(m-sulfonamido) benzyl	(m-hydroxy) benzyl	benzyl
5675	(m-sulfonamido) benzyl	(p-hydroxy) benzyl	benzyl
5676	(m-sulfonamido) benzyl	(m-hydroxymethyl) benzyl	benzyl
5677	(m-sulfonamido) benzyl	(p-hydroxymethyl) benzyl	benzyl
5678	(m-sulfonamido) benzyl	(m-carboxamido) benzyl	benzyl
5679	(m-sulfonamido) benzyl	(m- (N-methyl carboxamido)) benzyl	benzyl
5680	(m-sulfonamido) benzyl	(m-acetyl) benzyl	benzyl
5681	(m-sulfonamido) benzyl	(m-glycolyl) benzyl	benzyl
5682	(m-sulfonamido) benzyl	(m-sulfonamido) benzyl	benzyl
5683	(m-sulfonamido) benzyl	(m- (N-methylamino) benzyl	benzyl

5684	m-(N-ethylcarboxamido) benzyl	(m-hydroxy) benzyl	benzyl
5685	m-(N-ethylcarboxamido) benzyl	(p-hydroxy) benzyl	benzyl
5686	m-(N-ethylcarboxamido) benzyl	(m-hydroxymethyl) benzyl	benzyl
5687	m-(N-ethylcarboxamido) benzyl	(p-hydroxymethyl) benzyl	benzyl
5688	m-(N-ethylcarboxamido) benzyl	(m-carboxamido) benzyl	benzyl
5689	m-(N-ethylcarboxamido) benzyl	(m-(N-methyl carboxamido)) benzyl	benzyl
5690	m-(N-ethylcarboxamido) benzyl	(m-acetyl) benzyl	benzyl
5691	m-(N-ethylcarboxamido) benzyl	(m-glycolyl) benzyl	benzyl
5692	m-(N-ethylcarboxamido) benzyl	(m-sulfonamido) benzyl	benzyl
5693	m-(N-ethylcarboxamido) benzyl	(m-(N-methylamino) benzyl	benzyl
5694	3-pyridylmethyl	(m-hydroxy) benzyl	benzyl
5695	3-pyridylmethyl	(p-hydroxy) benzyl	benzyl
5696	3-pyridylmethyl	(m-hydroxymethyl) benzyl	benzyl
5697	3-pyridylmethyl	(p-hydroxymethyl) benzyl	benzyl
5698	3-pyridylmethyl	(m-carboxamido) benzyl	benzyl
5699	3-pyridylmethyl	(m-(N-methyl carboxamido)) benzyl	benzyl
5700	3-pyridylmethyl	(m-acetyl) benzyl	benzyl
5701	3-pyridylmethyl	(m-glycolyl) benzyl	benzyl
5702	3-pyridylmethyl	(m-sulfonamido) benzyl	benzyl
5703	3-pyridylmethyl	(m-(N-methylamino) benzyl	benzyl

5704	(m-carboxy)benzyl	(m-hydroxy)benzyl	(p-fluoro)benzyl
5705	(m-carboxy)benzyl	(p-hydroxy)benzyl	(p-fluoro)benzyl
5706	(m-carboxy)benzyl	(m-hydroxymethyl) benzyl	(p-fluoro)benzyl
5707	(m-carboxy)benzyl	(p-hydroxymethyl) benzyl	(p-fluoro)benzyl
5708	(m-carboxy)benzyl	(m-carboxamido) benzyl	(p-fluoro)benzyl
5709	(m-carboxy)benzyl	(m-(N-methyl carboxamido))benzyl	(p-fluoro)benzyl
5710	(m-carboxy)benzyl	(m-acetyl)benzyl	(p-fluoro)benzyl
5711	(m-carboxy)benzyl	(m-glycolyl)benzyl	(p-fluoro)benzyl
5712	(m-carboxy)benzyl	(m-sulfonamido) benzyl	(p-fluoro)benzyl
5713	(m-carboxy)benzyl	(m-(N-methylamino) benzyl	(p-fluoro)benzyl
5714	(m-carboxamido)benzyl	(m-hydroxy)benzyl	(p-fluoro)benzyl
5715	(m-carboxamido)benzyl	(p-hydroxy)benzyl	(p-fluoro)benzyl
5716	(m-carboxamido)benzyl	(m-hydroxymethyl) benzyl	(p-fluoro)benzyl
5717	(m-carboxamido)benzyl	(p-hydroxymethyl) benzyl	(p-fluoro)benzyl
5718	(m-carboxamido)benzyl	(m-carboxamido) benzyl	(p-fluoro)benzyl
5719	(m-carboxamido)benzyl	(m-(N-methyl carboxamido))benzyl	(p-fluoro)benzyl
5720	(m-carboxamido)benzyl	(m-acetyl)benzyl	(p-fluoro)benzyl
5721	(m-carboxamido)benzyl	(m-glycolyl)benzyl	(p-fluoro)benzyl
5722	(m-carboxamido)benzyl	(m-sulfonamido) benzyl	(p-fluoro)benzyl
5723	(m-carboxamido)benzyl	(m-(N-methylamino) benzyl	(p-fluoro)benzyl
5724	(m-(N-methyl carboxamido))benzyl	(m-hydroxy)benzyl	(p-fluoro)benzyl
5725	(m-(N-methyl carboxamido))benzyl	(p-hydroxy)benzyl	(p-fluoro)benzyl

5726	(m- (N-methyl carboxamido)) benzyl	(m-hydroxymethyl) benzyl	(p-fluoro) benzyl
5727	(m- (N-methyl carboxamido)) benzyl	(p-hydroxymethyl) benzyl	(p-fluoro) benzyl
5728	(m- (N-methyl carboxamido)) benzyl	(m-carboxamido) benzyl	(p-fluoro) benzyl
5729	(m- (N-methyl carboxamido)) benzyl	(m- (N-methyl carboxamido)) benzyl	(p-fluoro) benzyl
5730	(m- (N-methyl carboxamido)) benzyl	(m-acetyl) benzyl	(p-fluoro) benzyl
5731	(m- (N-methyl carboxamido)) benzyl	(m-glycolyl) benzyl	(p-fluoro) benzyl
5732	(m- (N-methyl carboxamido)) benzyl	(m-sulfonamido) benzyl	(p-fluoro) benzyl
5733	(m- (N-methyl carboxamido)) benzyl	(m- (N-methylamino) benzyl	(p-fluoro) benzyl
5734	(m-sulfonamido) benzyl	(m-hydroxy) benzyl	(p-fluoro) benzyl
5735	(m-sulfonamido) benzyl	(p-hydroxy) benzyl	(p-fluoro) benzyl
5736	(m-sulfonamido) benzyl	(m-hydroxymethyl) benzyl	(p-fluoro) benzyl
5737	(m-sulfonamido) benzyl	(p-hydroxymethyl) benzyl	(p-fluoro) benzyl
5738	(m-sulfonamido) benzyl	(m-carboxamido) benzyl	(p-fluoro) benzyl
5739	(m-sulfonamido) benzyl	(m- (N-methyl carboxamido)) benzyl	(p-fluoro) benzyl
5740	(m-sulfonamido) benzyl	(m-acetyl) benzyl	(p-fluoro) benzyl
5741	(m-sulfonamido) benzyl	(m-glycolyl) benzyl	(p-fluoro) benzyl
5742	(m-sulfonamido) benzyl	(m-sulfonamido) benzyl	(p-fluoro) benzyl
5743	(m-sulfonamido) benzyl	(m- (N-methylamino) benzyl	(p-fluoro) benzyl
5744	m- (N-ethylcarboxamido) benzyl	(m-hydroxy) benzyl	(p-fluoro) benzyl
5745	m- (N-ethylcarboxamido) benzyl	(p-hydroxy) benzyl	(p-fluoro) benzyl

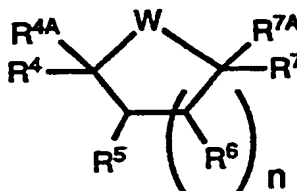
5746	m-(N-ethylcarboxamido) benzyl	(m-hydroxymethyl) benzyl	(p-fluoro)benzyl
5747	m-(N-ethylcarboxamido) benzyl	(p-hydroxymethyl) benzyl	(p-fluoro)benzyl
5748	m-(N-ethylcarboxamido) benzyl	(m-carboxamido) benzyl	(p-fluoro)benzyl
5749	m-(N-ethylcarboxamido) benzyl	(m-(N-methyl carboxamido)) benzyl	(p-fluoro)benzyl
5750	m-(N-ethylcarboxamido) benzyl	(m-acetyl)benzyl	(p-fluoro)benzyl
5751	m-(N-ethylcarboxamido) benzyl	(m-glycolyl)benzyl	(p-fluoro)benzyl
5752	m-(N-ethylcarboxamido) benzyl	(m-sulfonamido) benzyl	(p-fluoro)benzyl
5753	m-(N-ethylcarboxamido) benzyl	(m-(N-methylamino) benzyl	(p-fluoro)benzyl
5754	3-pyridylmethyl	(m-hydroxy)benzyl	(p-fluoro)benzyl
5755	3-pyridylmethyl	(p-hydroxy)benzyl	(p-fluoro)benzyl
5756	3-pyridylmethyl	(m-hydroxymethyl) benzyl	(p-fluoro)benzyl
5757	3-pyridylmethyl	(p-hydroxymethyl) benzyl	(p-fluoro)benzyl
5758	3-pyridylmethyl	(m-carboxamido) benzyl	(p-fluoro)benzyl
5759	3-pyridylmethyl	(m-(N-methyl carboxamido)) benzyl	(p-fluoro)benzyl
5760	3-pyridylmethyl	(m-acetyl)benzyl	(p-fluoro)benzyl
5761	3-pyridylmethyl	(m-glycolyl)benzyl	(p-fluoro)benzyl
5762	3-pyridylmethyl	(m-sulfonamido) benzyl	(p-fluoro)benzyl
5763	3-pyridylmethyl	(m-(N-methylamino) benzyl	(p-fluoro)benzyl

CLAIMS

WHAT IS CLAIMED IS:

5

1. A compound of the formula (I):



(I)

10

or a pharmaceutically acceptable salt or prodrug form thereof wherein:

15 R⁴ and R⁷ are independently selected from the following groups:

- hydrogen;
- C₁-C₈ alkyl substituted with 0-3 R¹¹;
- C₂-C₈ alkenyl substituted with 0-3 R¹¹;
- 20 C₂-C₈ alkynyl substituted with 0-3 R¹¹;
- C₃-C₈ cycloalkyl substituted with 0-3 R¹¹;
- C₆-C₁₀ bicycloalkyl substituted with 0-3 R¹¹;
- aryl substituted with 0-3 R¹²;
- a C₆-C₁₄ carbocyclic residue substituted with 0-3 R¹²;
- 25 a heterocyclic ring system substituted with 0-2 R¹², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

30

R^{4A} and R^{7A} are independently selected from the following groups:

- 5 hydrogen;
 C₁-C₄ alkyl substituted with halogen or C₁-C₂
 alkoxy;
 benzyl substituted with halogen or C₁-C₂ alkoxy;

10 R⁴ and R^{4A} can alternatively join to form a 5-7 membered carbocyclic ring substituted with 0-2 R¹²;

R⁷ and R^{7A} can alternatively join to form a 5-7 membered carbocyclic ring substituted with 0-2 R¹²;

15 n is 0, 1, or 2;

R⁵ is selected from fluoro, difluoro, =O, C₁-C₃ alkyl or -OR²⁰;

20 R⁶, when n is 1, is selected from: hydrogen, =O, fluoro, difluoro, C₁-C₃ alkyl or -OR²¹;

R⁶, when n is 2, is independently selected from: hydrogen, =O, fluoro, difluoro, C₁-C₃ alkyl or -OR²¹;

25

R⁵ and R⁶ can alternatively join to form an epoxide ring; -OCH₂SCH₂O-; -OS(=O)O-; -OC(=O)O-; -OCH₂O-; -OC(=S)O-; -OC(=O)C(=O)O-; -OC(CH₃)₂O-; -OC(OCH₃)(CH₂CH₂CH₃)O-; or any group that, when
30 administered to a mammalian subject, cleaves to form a free dihydroxyl;

R²⁰ and R²¹ are independently selected from:

35 hydrogen;

C₁-C₆ alkyl substituted with 0-3 R¹¹;
C₃-C₆ alkoxyalkyl substituted with 0-3 R¹¹;
C₁-C₆ alkylcarbonyl substituted with 0-3 R¹¹;
C₁-C₆ alkoxycarbonyl substituted with 0-3 R¹¹;
5 benzoyl substituted with 0-3 R¹²;
phenoxycarbonyl substituted with 0-3 R¹²;
phenylaminocarbonyl substituted with 0-3 R¹²; or
any group that, when administered to a mammalian
subject, cleaves to form a free hydroxyl;

10

R¹¹ is selected from one or more of the following:

keto, halogen, cyano, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴,
-CO₂R¹³, -OC(=O)R¹³, -OR¹³, C₂-C₆ alkoxyalkyl,
15 -S(O)_mR¹³, -NHC(=NH)NHR¹³, -C(=NH)NHR¹³,
-C(=O)NR¹³R¹⁴, -NR¹⁴C(=O)R¹³, =NOR¹⁴,
-NR¹⁴C(=O)OR¹⁴, -OC(=O)NR¹³R¹⁴, -NR¹³C(=O)NR¹³R¹⁴,
-NR¹⁴SO₂NR¹³R¹⁴, -NR¹⁴SO₂R¹³, -SO₂NR¹³R¹⁴, C₁-C₄
alkyl, C₂-C₄ alkenyl, C₃-C₆ cycloalkyl, C₃-C₆
20 cycloalkylmethyl;

1-3 amino acids, linked together via amide bonds
and linked to R⁴ or R⁷ via the amine or carboxylate
terminus;

25

a C₅-C₁₄ carbocyclic residue substituted with 0-3
R¹²;

aryl substituted with 0-3 R¹²; or

30

a heterocyclic ring system substituted with 0-2
R¹², composed of 5 to 10 atoms including at least
one nitrogen, oxygen or sulfur atom;

35

R¹², when a substituent on carbon, is selected from one or more of the following:

5 phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
halogen, hydroxy, nitro, cyano, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, C₇-C₁₀ arylalkyl,
C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide,
oxime, boronic acid, sulfonamide, formyl, C₃-C₆
10 cycloalkoxy, -OR¹³, C₁-C₄ alkyl substituted with
-NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄
hydroxyalkyl, methylenedioxy, ethylenedioxy, C₁-C₄
haloalkyl, C₁-C₄ haloalkoxy, C₁-C₄ alkoxycarbonyl,
C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, C₁-C₄
alkylcarbonylamino, -S(O)_mR¹³, -SO₂NR¹³R¹⁴,
15 -NHSO₂R¹⁴, -OCH₂CO₂H, 2-(1-morpholino)ethoxy; or

a 5- or 6-membered heterocyclic ring containing from
1 to 4 heteroatoms selected from oxygen, nitrogen or
sulfur;

20 or R¹² may be a 3- or 4- carbon chain attached to
adjacent carbons on the ring to form a fused 5- or 6-
membered ring, said 5- or 6- membered ring being
optionally substituted on the aliphatic carbons with
25 halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, hydroxy, or
-NR¹³R¹⁴; or, when R¹² is attached to a saturated
carbon atom, it may be carbonyl or thiocarbonyl;

30 R¹², when a substituent on nitrogen, is selected from
one or more of the following:

phenyl, benzyl, phenethyl, hydroxy, C₁-C₄
hydroxyalkyl, C₁-C₄ alkoxy, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, -CH₂NR¹³R¹⁴,
35 -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ haloalkyl, C₁-C₄

alkoxycarbonyl, $-\text{CO}_2\text{H}$, $\text{C}_1\text{-C}_4$ alkylcarbonyloxy,
 $\text{C}_1\text{-C}_4$ alkylcarbonyl;

5 R^{13} is H, phenyl, benzyl, $\text{C}_1\text{-C}_6$ alkyl, or $\text{C}_3\text{-C}_6$
alkoxyalkyl;

R^{14} is OH, H, $\text{C}_1\text{-C}_4$ alkyl, or benzyl;

10 R^{13} and R^{14} can alternatively join to form $-(\text{CH}_2)_4-$,
 $-(\text{CH}_2)_5-$, $-\text{CH}_2\text{CH}_2\text{N}(\text{R}^{15})\text{CH}_2\text{CH}_2-$, or $-\text{CH}_2\text{CH}_2\text{OCH}_2\text{CH}_2-$;

R^{15} is H or CH_3 ;

m is 0, 1 or 2;

15

W is selected from:

20 $-\text{N}(\text{R}^{22})\text{C}(=\text{Z})\text{N}(\text{R}^{23})-$;
 $-\text{N}(\text{R}^{22})\text{S}(=\text{O})\text{N}(\text{R}^{23})-$;
 $-\text{OC}(=\text{Z})\text{O}-$;
 $-\text{N}(\text{R}^{22})\text{C}(=\text{Z})\text{O}-$;
 $-\text{C}(\text{R}^{25})(\text{R}^{26})\text{C}(=\text{Z})\text{C}(\text{R}^{27})(\text{R}^{28})-$;
 $-\text{N}(\text{R}^{22})\text{C}(=\text{Z})\text{C}(\text{R}^{27})(\text{R}^{28})-$;
 $-\text{C}(\text{R}^{25})(\text{R}^{26})\text{C}(=\text{Z})\text{O}-$;
 $-\text{N}(\text{R}^{22})\text{C}(=\text{O})\text{C}(=\text{O})\text{N}(\text{R}^{23})-$;
25 $-\text{C}(\text{R}^{25})(\text{R}^{26})\text{C}(\text{F}_2)\text{C}(\text{R}^{27})(\text{R}^{28})-$;
 $-\text{C}(\text{R}^{25})(\text{R}^{26})\text{N}(\text{CH}_3)(\text{O})\text{C}(\text{R}^{27})(\text{R}^{28})-$;
 $-\text{C}(\text{R}^{25})(\text{R}^{26})\text{N}(\text{OR}^{29})\text{C}(\text{R}^{27})(\text{R}^{28})-$;
 $-\text{C}(\text{R}^{25})(\text{R}^{26})\text{C}(=\text{Z})\text{S}-$;

30 wherein:

Z is O, S, or NR^{24} ;

35 R^{22} and R^{23} are independently selected from the
following:

- hydrogen;
C₁-C₈ alkyl substituted with 0-3 R³¹;
C₂-C₈ alkenyl substituted with 0-3 R³¹;
C₂-C₈ alkynyl substituted with 0-3 R³¹;
5 C₃-C₈ cycloalkyl substituted with 0-3 R³¹;
C₆-C₁₀ bicycloalkyl substituted with 0-3 R³¹;
aryl substituted with 0-3 R³²;
a C₆-C₁₄ carbocyclic residue substituted with 0-3
R³²;
10 a heterocyclic ring system substituted with 0-2
R³², composed of 5 to 10 atoms including at least
one nitrogen, oxygen or sulfur atom;

R²⁴ is selected from: hydroxy; amino; C₁-C₄ alkyl; C₁-C₄
15 alkoxy; C₁-C₄ aminoalkyl; cyano; nitro; benzyloxy;

R²⁵ and R²⁷ are independently selected from the
following:

- 20 hydrogen;
C₁-C₈ alkyl substituted with 0-3 R³¹;
C₂-C₈ alkenyl substituted with 0-3 R³¹;
C₂-C₈ alkynyl substituted with 0-3 R³¹;
C₃-C₈ cycloalkyl substituted with 0-3 R³¹;
25 C₆-C₁₀ bicycloalkyl substituted with 0-3 R³¹;
aryl substituted with 0-3 R³²;
a C₆-C₁₄ carbocyclic residue substituted with 0-3
R³²;
a heterocyclic ring system substituted with 0-2
30 R³², composed of 5 to 10 atoms including at least
one nitrogen, oxygen or sulfur atom;

R²⁶ and R²⁸ are independently selected from:

- 35 hydrogen;

C₁-C₄ alkyl substituted with halogen or C₁-C₂
alkoxy;
benzyl substituted with halogen or C₁-C₂ alkoxy;

5 R²⁹ is selected from:

hydrogen;
C₁-C₄ alkyl substituted with halogen or C₁-C₂
alkoxy;
10 benzyl substituted with halogen or C₁-C₂ alkoxy;

alternatively, R²², R²⁵, or R²⁶, independently, can join
with R⁴ or R^{4A} to form a five- or six-membered fused
15 heterocyclic or carbocyclic ring substituted with 0-2
R¹²; and

alternatively, R²³, R²⁷, or R²⁸, independently, can join
with R⁷ or R^{7A} to form a five- or six-membered fused
20 heterocyclic or carbocyclic ring substituted with 0-2
R¹²; and

alternatively, W can join with R⁵ or R⁶ to form a three-
to seven-membered fused heterocyclic or carbocyclic ring
25 substituted with 0-2 R¹²;

R³¹ is selected from one or more of the following:

keto, halogen, cyano, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴,
30 -CO₂R¹³,
-OC(=O)R¹³, -OR¹³, C₂-C₆ alkoxyalkyl, -S(O)_mR¹³,
-NHC(=NH)NHR¹³, -C(=NH)NHR¹³, -C(=O)NR¹³R¹⁴,
-NR¹⁴C(=O)R¹³, =NOR¹⁴, -NR¹⁴C(=O)OR¹⁴,
-OC(=O)NR¹³R¹⁴, -NR¹³C(=O)NR¹³R¹⁴, -NR¹⁴SO₂NR¹³R¹⁴,
35 -NR¹⁴SO₂R¹³, -SO₂NR¹³R¹⁴, C₁-C₄ alkyl, C₂-C₄
alkenyl, C₃-C₆ cycloalkyl, C₃-C₆ cycloalkylmethyl;

1-3 amino acids, linked together via amide bonds and linked to R⁴ or R⁷ via the amine or carboxylate terminus;

5

a C₅-C₁₄ carbocyclic residue substituted with 0-3 R³²;

aryl substituted with 0-3 R³²; or

10

a heterocyclic ring system substituted with 0-2 R³², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

15 R³², when a substituent on carbon, is selected from one or more of the following:

phenyl, benzyl, phenethyl, phenoxy, benzyloxy, halogen, hydroxy, nitro, cyano, C₁-C₄ alkyl, C₃-C₆ cycloalkyl, C₃-C₆ cycloalkylmethyl, C₇-C₁₀ arylalkyl, C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide, oxime, boronic acid, sulfonamide, formyl, C₃-C₆ cycloalkoxy, -OR¹³, C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ hydroxyalkyl, methylenedioxy, ethylenedioxy, C₁-C₄ haloalkyl, C₁-C₄ haloalkoxy, C₁-C₄ alkoxycarbonyl, C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, C₁-C₄ alkylcarbonylamino, -S(O)_mR¹³, -SO₂NR¹³R¹⁴, -NHSO₂R¹⁴, -OCH₂CO₂H, 2-(1-morpholino)ethoxy, -C(R¹⁴)=N(OR¹⁴); or

25
30

a 5- or 6-membered heterocyclic ring containing from 1 to 4 heteroatoms selected from oxygen, nitrogen or sulfur;

35

or R³² may be a 3- or 4- carbon chain attached to adjacent carbons on the ring to form a fused 5- or 6-membered ring, said 5- or 6- membered ring being optionally substituted on the aliphatic carbons with
5 halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, hydroxy, or -NR¹³R¹⁴; or, when R³² is attached to a saturated carbon atom, it may be carbonyl or thiocarbonyl;

R³², when a substituent on nitrogen, is selected from
10 one or more of the following:

phenyl, benzyl, phenethyl, hydroxy, C₁-C₄ hydroxyalkyl, C₁-C₄ alkoxy, C₁-C₄ alkyl, C₃-C₆ cycloalkyl, C₃-C₆ cycloalkylmethyl, -CH₂NR¹³R¹⁴,
15 -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ haloalkyl, C₁-C₄ alkoxycarbonyl, -CO₂H, C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, -C(R¹⁴)=N(OR¹⁴);

provided that:
20

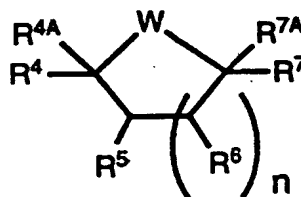
R⁴, R^{4A}, R⁷, and, R^{7A} are not all hydrogen;

when W is -OC(=Z)O-, R⁴ and R⁷ are not hydrogen;

25 when R⁴ and R^{4A} are both hydrogen, at least one of the following is not hydrogen: R²², R²³, R²⁵, R²⁶, R²⁷ and R²⁸.

30

2. A compound of Claim 1 of formula (I):



(I)

or a pharmaceutically acceptable salt or prodrug form
5 thereof wherein:

R⁴ and R⁷ are independently selected from the following
groups:

10 hydrogen;

C₁-C₄ alkyl substituted with 0-3 R¹¹;

C₃-C₄ alkenyl substituted with 0-3 R¹¹;

C₃-C₄ alkynyl substituted with 0-3 R¹¹;

15 R^{4A} and R^{7A} are hydrogen;

n is 0 or 1;

R⁵ is selected from fluoro, difluoro, =O, or -OR²⁰;

20

R⁶ is selected from: hydrogen, =O, fluoro, difluoro, or
-OR²¹;

25 R⁵ and R⁶ can alternatively join to form an epoxide
ring; -OCH₂SCH₂O-; -OS(=O)O-; -OC(=O)O-; -OCH₂O-;
-OC(=S)O-; -OC(=O)C(=O)O-; -OC(CH₃)₂O-;
-OC(OCH₃)(CH₂CH₂CH₃)O-; or any group that, when
administered to a mammalian subject, cleaves to
form a free dihydroxyl;

30

R²⁰ and R²¹ are independently selected from:

hydrogen;
C₁-C₆ alkylcarbonyl;
5 C₁-C₆ alkoxy carbonyl;
benzoyl; or
any group that, when administered to a mammalian
subject, cleaves to form a free hydroxyl;

10

R¹¹ is selected from one or more of the following:

keto, halogen, cyano, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴,
-CO₂R¹³,
15 -OC(=O)R¹³, -OR¹³, C₂-C₄ alkoxyalkyl, -S(O)_mR¹³,
C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆ cycloalkyl;

a C₅-C₁₄ carbocyclic residue substituted with 0-3
R¹²;

20

aryl substituted with 0-3 R¹²; or

a heterocyclic ring system substituted with 0-2
R¹², composed of 5 to 10 atoms including at least
25 one nitrogen, oxygen or sulfur atom;

25

R¹², when a substituent on carbon, is selected from one
or more of the following:

30

phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
halogen, hydroxy, nitro, cyano, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, C₇-C₁₀ arylalkyl,
C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide,
35 oxime, boronic acid, sulfonamide, formyl, C₃-C₆

cycloalkoxy, $-OR^{13}$, C₁-C₄ alkyl substituted with
-NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄
hydroxyalkyl, methylenedioxy, ethylenedioxy, C₁-C₄
haloalkyl, C₁-C₄ haloalkoxy, C₁-C₄ alkoxycarbonyl,
5 C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, C₁-C₄
alkylcarbonylamino, $-S(O)_mR^{13}$,
-SO₂NR¹³R¹⁴, -NHSO₂R¹⁴; or

10 a 5- or 6-membered heterocyclic ring containing from
1 to 4 heteroatoms selected from oxygen, nitrogen or
sulfur;

or R¹² may be a 3- or 4- carbon chain attached to
adjacent carbons on the ring to form a fused 5- or 6-
15 membered ring, said 5- or 6- membered ring being
optionally substituted on the aliphatic carbons with
halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, hydroxy, or
-NR¹³R¹⁴; or, when R¹² is attached to a saturated
carbon atom, it may be carbonyl or thiocarbonyl;

20 R¹², when a substituent on nitrogen, is selected from
one or more of the following:

phenyl, benzyl, phenethyl, hydroxy, C₁-C₄
25 hydroxyalkyl, C₁-C₄ alkoxy, C₁-C₄ alkyl, C₃-C₆
cycloalkyl, C₃-C₆ cycloalkylmethyl, -CH₂NR¹³R¹⁴,
-NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ haloalkyl, C₁-C₄
alkoxycarbonyl, C₁-C₄ alkylcarbonyloxy, C₁-C₄
alkylcarbonyl, -CO₂H;

30 R¹³ is H, C₁-C₆ alkyl, or C₃-C₆ alkoxyalkyl;

R¹⁴ is OH, H, C₁-C₄ alkyl, or benzyl;

35 R¹³ and R¹⁴ can alternatively join to form $-(CH_2)_4-$,
 $-(CH_2)_5-$, $-CH_2CH_2N(R^{15})CH_2CH_2-$, or $-CH_2CH_2OCH_2CH_2-$;

R¹⁵ is H or CH₃;

m is 0, 1 or 2;

5

W is selected from:

- 10
- N(R²²)C(=Z)N(R²³)-;
 - N(R²²)C(=Z)O-;
 - C(R²⁵)(R²⁶)C(=Z)C(R²⁷)(R²⁸)-;
 - N(R²²)C(=Z)C(R²⁷)(R²⁸)-;
 - C(R²⁵)(R²⁶)C(=Z)O-;
 - N(R²²)C(=O)C(=O)N(R²³)-;
 - C(R²⁵)(R²⁶)C(F₂)C(R²⁷)(R²⁸)-;

15

wherein:

Z is O, S, N-CN, N-OH, N-OCH₃;

- 20
- R²² and R²³ are independently selected from the following:

- 25
- hydrogen;
 - C₁-C₈ alkyl substituted with 0-3 R³¹;
 - C₃-C₈ alkenyl substituted with 0-3 R³¹;
 - C₃-C₈ alkynyl substituted with 0-3 R³¹;
 - C₃-C₆ cycloalkyl substituted with 0-3 R³¹;

- 30
- R²⁵ and R²⁷ are independently selected from the following:

- 35
- hydrogen;
 - C₁-C₈ alkyl substituted with 0-3 R³¹;
 - C₂-C₈ alkenyl substituted with 0-3 R³¹;
 - C₃-C₈ alkynyl substituted with 0-3 R³¹;

R²⁶ and R²⁸ are hydrogen;

R³¹ is selected from one or more of the following:

- 5 keto, halogen, cyano, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴,
 -CO₂R¹³, -OC(=O)R¹³, -OR¹³, C₂-C₄ alkoxyalkyl,
 -S(O)_mR¹³, C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆
 cycloalkyl;
- 10 a C₅-C₁₄ carbocyclic residue substituted with 0-3
 R¹²;
- aryl substituted with 0-3 R³²; or
- 15 a heterocyclic ring system substituted with 0-2
 R³², composed of 5 to 10 atoms including at least
 one nitrogen, oxygen or sulfur atom;
- 20 R³², when a substituent on carbon, is selected from one
 or more of the following:
- phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
 halogen, hydroxy, nitro, cyano, C₁-C₄ alkyl, C₃-C₆
25 cycloalkyl, C₃-C₆ cycloalkylmethyl, C₇-C₁₀ arylalkyl,
 C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide,
 oxime, boronic acid, sulfonamide, formyl, C₃-C₆
 cycloalkoxy, -OR¹³, C₁-C₄ alkyl substituted with
 -NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄
30 hydroxyalkyl, methylenedioxy, ethylenedioxy, C₁-C₄
 haloalkyl, C₁-C₄ haloalkoxy, C₁-C₄ alkoxycarbonyl,
 C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, C₁-C₄
 alkylcarbonylamino, -S(O)_mR¹³,
 -SO₂NR¹³R¹⁴, -NHSO₂R¹⁴, -C(R¹⁴)=N(OR¹⁴); or
35

a 5- or 6-membered heterocyclic ring containing from 1 to 4 heteroatoms selected from oxygen, nitrogen or sulfur;

5 or R³² may be a 3- or 4- carbon chain attached to adjacent carbons on the ring to form a fused 5- or 6-membered ring, said 5- or 6- membered ring being optionally substituted on the aliphatic carbons with halogen, C₁-C₄ alkyl, C₁-C₄ alkoxy, hydroxy, or
10 -NR¹³R¹⁴; or, when R³² is attached to a saturated carbon atom, it may be carbonyl or thiocarbonyl;

R³², when a substituent on nitrogen, is selected from one or more of the following:

15 phenyl, benzyl, phenethyl, hydroxy, C₁-C₄ hydroxyalkyl, C₁-C₄ alkoxy, C₁-C₄ alkyl, C₃-C₆ cycloalkyl, C₃-C₆ cycloalkylmethyl, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, C₂-C₆ alkoxyalkyl, C₁-C₄ haloalkyl, C₁-C₄
20 alkoxy carbonyl, C₁-C₄ alkylcarbonyloxy, C₁-C₄ alkylcarbonyl, -CO₂H, -C(R¹⁴)=N(OR¹⁴);

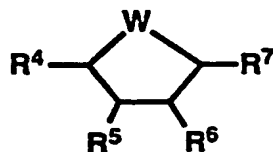
provided that:

25 R⁴, R^{4A}, R⁷, and R^{7A} are not all hydrogen;

when W is -OC(=Z)O-, R⁴ and R⁷ are not hydrogen;

30 when R⁴ and R^{4A} are both hydrogen, at least one of the following is not hydrogen: R²², R²³, R²⁵, R²⁶, R²⁷ and R²⁸.

35 3. A compound of Claim 1 of formula (II):



5 or a pharmaceutically acceptable salt or prodrug form thereof wherein:

R⁴ and R⁷ are independently selected from the following groups:

10

hydrogen;

C₁-C₄ alkyl substituted with 0-3 R¹¹;

C₃-C₄ alkenyl substituted with 0-3 R¹¹;

15 R⁵ is -OR²⁰;

R⁶ is hydrogen or -OR²¹;

20 R²⁰ and R²¹ are independently hydrogen or any group that, when administered to a mammalian subject, cleaves to form a free hydroxyl;

R¹¹ is selected from one or more of the following:

25 keto, halogen, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, -OR¹³, C₂-C₄ alkoxyalkyl, C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆ cycloalkyl;

aryl substituted with 0-3 R¹²; or

30

a heterocyclic ring system substituted with 0-2 R¹², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

R¹², when a substituent on carbon, is selected from one or more of the following:

- 5 phenyl, benzyl, phenethyl, phenoxy, benzyloxy, halogen, C₁-C₄ alkyl, C₇-C₁₀ arylalkyl, C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide, oxime, boronic acid, sulfonamide, formyl, C₃-C₆ cycloalkoxy, -OR¹³,
10 C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴, methylenedioxy, C₁-C₄ haloalkyl, C₁-C₄ alkylcarbonyl, C₁-C₄ alkylcarbonylamino, hydroxy, hydroxymethyl; or
15 a 5- or 6-membered heterocyclic ring containing from 1 to 4 heteroatoms selected from oxygen, nitrogen or sulfur;

R¹², when a substituent on nitrogen, is selected from benzyl or methyl;

- 20 R¹³ is H, C₁-C₂ alkyl, or C₃-C₆ alkoxyalkyl;

R¹⁴ is OH, H or C₁-C₂ alkyl;

- 25 R¹³ and R¹⁴ can alternatively join to form -(CH₂)₄-, -(CH₂)₅-, -CH₂CH₂N(R¹⁵)CH₂CH₂-, or -CH₂CH₂OCH₂CH₂-;

W is selected from:

- 30 -N(R²²)C(=Z)N(R²³)-;
-C(R²⁵)(R²⁶)C(=Z)C(R²⁷)(R²⁸)-;
-N(R²²)C(=Z)C(R²⁷)(R²⁸)-;
-C(R²⁵)(R²⁶)C(=Z)O-;

- 35 wherein:

Z is O, S, or N-CN;

R²² and R²³ are independently selected from the following:

5

hydrogen;

C₁-C₄ alkyl substituted with 0-3 R³¹;

C₃-C₄ alkenyl substituted with 0-3 R³¹;

10 R²⁵ and R²⁷ are independently selected from the following:

hydrogen;

C₁-C₄ alkyl substituted with 0-3 R³¹;

15 C₃-C₄ alkenyl substituted with 0-3 R³¹;

R²⁶ and R²⁸ are hydrogen;

R³¹ is selected from one or more of the following:

20

keto, halogen, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, -OR¹³, C₂-C₄ alkoxyalkyl, C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆ cycloalkyl;

25 aryl substituted with 0-3 R³²; or

a heterocyclic ring system substituted with 0-2 R³², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

30

R³², when a substituent on carbon, is selected from one or more of the following:

phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
35 halogen, C₁-C₄ alkyl, C₇-C₁₀ arylalkyl, C₁-C₄ alkoxy, -CO₂H, hydroxamic acid, hydrazide, oxime, boronic

acid, sulfonamide, formyl, C₃-C₆ cycloalkoxy, -OR¹³,
C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴,
methylenedioxy, C₁-C₄ haloalkyl, C₁-C₄ alkylcarbonyl,
C₁-C₄ alkylcarbonylamino, hydroxy, hydroxymethyl,
5 -C(R¹⁴)=N(OR¹⁴); or

a 5- or 6-membered heterocyclic ring containing from
1 to 4 heteroatoms selected from oxygen, nitrogen or
sulfur;

10 R³², when a substituent on nitrogen, is selected from
benzyl or methyl;

provided that:

15 when R⁴ is hydrogen, R⁷ is not hydrogen;

when R⁴ is hydrogen, at least one of the following
is not hydrogen: R²², R²³, R²⁵, R²⁶, R²⁷ and R²⁸.

20

4. A compound of Claim 3 wherein:

R⁴ and R⁷ are independently selected from the following
25 groups:

hydrogen;
C₁-C₃ alkyl substituted with 0-1 R¹¹;

30 R⁵ is -OR²⁰;

R⁶ is hydrogen or -OR²¹;

R²⁰ and R²¹ are independently hydrogen or any group
35 that, when administered to a mammalian subject, cleaves
to form a free hydroxyl;

R¹¹ is selected from one or more of the following:

halogen, -OR¹³, C₁-C₄ alkyl, C₃-C₅ cycloalkyl;

5

aryl substituted with 0-2 R¹²; or

a heterocyclic ring system chosen from pyridyl,
pyrimidinyl, triazinyl, furanyl, thienyl, pyrrolyl,
10 pyrazolyl, imidazolyl, tetrazolyl, benzofuranyl,
indolyl, quinolinyl, isoquinolinyl;

R¹², when a substituent on carbon, is selected from one
or more of the following:

15

benzyloxy, halogen, methyl, C₁-C₄ alkoxy, CF₃,
2-(1-morpholino)ethoxy, -CO₂H, hydroxamic acid,
hydrazide, oxime, cyano, boronic acid, sulfonamide,
formyl, C₃-C₆ cycloalkoxy, C₁-C₄ alkyl substituted
20 with -NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl; or

R¹², when a substituent on nitrogen, is methyl;

R¹³ is H or methyl;

25

R¹⁴ is OH, H or methyl;

R¹³ and R¹⁴ can alternatively join to form -(CH₂)₄-,
-(CH₂)₅-, -CH₂CH₂N(R¹⁵)CH₂CH₂-, or -CH₂CH₂OCH₂CH₂-;

30

W is -N(R²²)C(=O)N(R²³)- or -N(R²²)C(=N-CN)N(R²³)-;

R²² and R²³ are independently selected from the
35 following:

hydrogen;
C₁-C₄ alkyl substituted with 0-1 R³¹;
C₃-C₄ alkenyl substituted with 0-1 R³¹;

5 R³¹ is selected from one or more of the following:

halogen, -OR¹³, C₁-C₄ alkyl, C₃-C₅ cycloalkyl;

aryl substituted with 0-2 R³²; or

10

a heterocyclic ring system chosen from pyridyl,
pyrimidinyl, triazinyl, furanyl, thienyl, pyrrolyl,
pyrazolyl, imidazolyl, tetrazolyl, benzofuranyl,
indolyl, quinolinyl, isoquinolinyl;

15

R³², when a substituent on carbon, is selected from one
or more of the following:

20 benzyloxy, halogen, methyl, C₁-C₄ alkoxy, CF₃,
2-(1-morpholino)ethoxy, -CO₂H, hydroxamic acid,
hydrazide, oxime, cyano, boronic acid, sulfonamide,
formyl, C₃-C₆ cycloalkoxy, C₁-C₄ alkyl substituted
with -NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl,
-C(R¹⁴)=N(OR¹⁴); or

25

R³², when a substituent on nitrogen, is methyl;

provided that:

30 when R⁴ is hydrogen, R⁷ is not hydrogen;

when R⁴ is hydrogen, at least one of the following
is not hydrogen: R²² and R²³.

35 5. A compound of Claim 3 wherein:

R⁴ and R⁷ are benzyl;

R⁵ is -OH;

5 R⁶ is hydrogen or -OH;

R¹³ is H or methyl;

R¹⁴ is H or methyl;

10

W is -N(R²²)C(=O)N(R²³)- or -N(R²²)C(=N-CN)N(R²³)-;

R²² and R²³ are independently selected from the following:

15

hydrogen;

C₁-C₄ alkyl substituted with 0-1 R³¹;

R³¹ is selected from one or more of the following:

20

C₃-C₅ cycloalkyl;

aryl substituted with 0-2 R³²; or

25

a heterocyclic ring system chosen from pyridyl, thienyl, quinolinyl, or isoquinolinyl;

R³², when a substituent on carbon, is selected from one or more of the following:

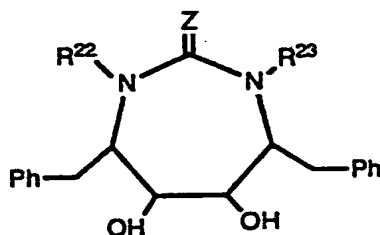
30

-CONH₂, -CO₂H, -CHO, -CH₂NHOH, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl, -C(R¹⁴)=N(OR¹⁴); or

R³², when a substituent on nitrogen, is methyl.

35

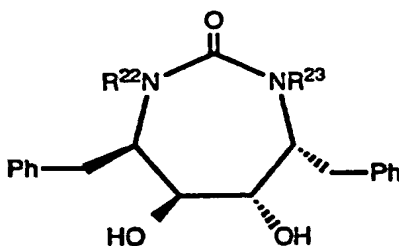
6. A compound of Claim 1 of formula:



5 wherein Z is O, S, or N-CN
 wherein R²² and R²³ are independently selected from the
 group consisting of:

hydrogen, allyl, propyl, cyclopropylmethyl,
 10 n-butyl, i-butyl, CH₂CH=CH(CH₃)₂, pyridylmethyl,
 methallyl, n-pentyl, i-pentyl, hexyl, benzyl,
 pyridylmethyl, isoprenyl, propargyl, picolinyl,
 methoxyethyl, cyclohexylmethyl, dimethyl-butyl,
 ethoxyethyl, methyl-oxazolinylmethyl,
 15 naphthylmethyl, methyloxazolinylmethyl,
 vinyloxyethyl, pentafluorobenzyl, quinolinylmethyl,
 carboxybenzyl, chloro-thienyl, picolinyl,
 benzyloxybenzyl, phenylbenzyl, adamantylethyl,
 cyclopropylmethoxybenzyl, ethoxybenzyl,
 20 hydroxybenzyl, hydroxymethylbenzyl, aminobenzyl,
 formylbenzyl, cyanobenzyl, cinnamyl,
 allyloxybenzyl, fluorobenzyl, cyclobutylmethyl,
 formaldoximebenzyl, cyclopentylmethyl, nitrobenzyl,
 nitrilobenzyl, carboxamidobenzyl,
 25 carbomethoxybenzyl, tetrazolylbenzyl and
 dimethylallyl.

7. A compound of Claim 1 of formula (IIa):
 30

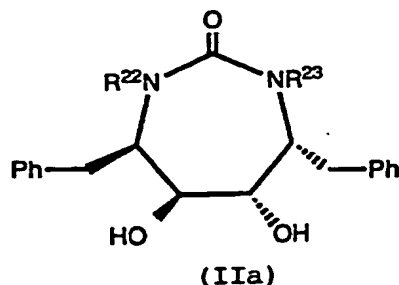


(IIa)

5 wherein R²² and R²³ are independently selected from the group consisting of:

hydrogen, allyl, propyl, cyclopropylmethyl,
 n-butyl, i-butyl, CH₂CH=CH(CH₃)₂, pyridylmethyl,
 10 methallyl, n-pentyl, i-pentyl, hexyl, benzyl,
 pyridylmethyl, isoprenyl, propargyl, picolinyl,
 methoxyethyl, cyclohexylmethyl, dimethyl-butyl,
 ethoxyethyl, methyl-oxazolinylmethyl,
 naphthylmethyl, methyloxazolinylmethyl,
 15 vinyloxyethyl, pentafluorobenzyl, quinolinylmethyl,
 carboxybenzyl, chloro-thienyl, picolinyl,
 benzyloxybenzyl, phenylbenzyl, adamantylethyl,
 cyclopropylmethoxybenzyl, ethoxybenzyl,
 hydroxybenzyl, hydroxymethylbenzyl, aminobenzyl,
 20 formylbenzyl, cyanobenzyl, cinnamyl,
 allyloxybenzyl, fluorobenzyl, cyclobutylmethyl,
 formaldoximebenzyl, cyclopentylmethyl, nitrobenzyl,
 nitrilobenzyl, carboxamidobenzyl,
 carbomethoxybenzyl, tetrazolylbenzyl,
 25 dimethylallyl, amidinobenzyl, and (boronic
 acid)benzyl.

8. A compound of Claim 1 of formula (IIa):
 30



which is selected from the group consisting of:

5

the compound of formula (IIa) wherein R²² is allyl and R²³ is allyl;

10 the compound of formula (IIa) wherein R²² is propyl and R²³ is propyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cyclopropylmethyl;

15 the compound of formula (IIa) wherein R²² is n-butyl and R²³ is n-butyl;

the compound of formula (IIa) wherein R²² is CH₂CH=CH(CH₃)₂ and R²³ is CH₂CH=CH(CH₃)₂;

20

the compound of formula (IIa) wherein R²² is i-pentyl and R²³ is i-pentyl;

25 the compound of formula (IIa) wherein R²² is 4-pyridylmethyl and R²³ is 4-pyridylmethyl;

the compound of formula (IIa) wherein R²² is 2-methallyl and R²³ is 2-methallyl;

30 the compound of formula (IIa) wherein R²² is n-pentyl and R²³ is n-pentyl;

the compound of formula (IIa) wherein R²² is i-butyl and R²³ is i-butyl;

5 the compound of formula (IIa) wherein R²² is benzyl and R²³ is benzyl;

the compound of formula (IIa) wherein R²² is 3-pyridylmethyl and R²³ is 3-pyridylmethyl;

10

the compound of formula (IIa) wherein R²² is allyl and R²³ is isoprenyl;

15 the compound of formula (IIa) wherein R²² is 3-propargyl and R²³ is 3-propargyl;

the compound of formula (IIa) wherein R²² is 2-picolinyl and R²³ is 2-picolinyl;

20 the compound of formula (IIa) wherein R²² is 2-methoxyethyl and R²³ is 2-methoxyethyl;

the compound of formula (IIa) wherein R²² is cyclohexylmethyl and R²³ is cyclohexylmethyl;

25

the compound of formula (IIa) wherein R²² is 3,3-dimethyl-1-butyl and R²³ is 3,3-dimethyl-1-butyl;

30 the compound of formula (IIa) wherein R²² is 2-ethoxyethyl and R²³ is 2-ethoxyethyl;

the compound of formula (IIa) wherein R²² is 3-methyl-5-oxazolinylmethyl and R²³ is hydrogen;

35 the compound of formula (IIa) wherein R²² is 1-naphthylmethyl and R²³ is 1-naphthylmethyl;

the compound of formula (IIa) wherein R²² is
3-methyloxazolinylmethyl and R²³ is
3-methyloxazolinylmethyl;

5

the compound of formula (IIa) wherein R²² is
2-vinyloxyethyl and R²³ is 2-vinyloxyethyl;

the compound of formula (IIa) wherein R²² is
10 2,3,4,5,6-pentafluorobenzyl and R²³ is
2,3,4,5,6-pentafluorobenzyl;

15

the compound of formula (IIa) wherein R²² is benzyl
and R²³ is 2-quinolinylmethyl;

the compound of formula (IIa) wherein R²² is
4-carboxybenzyl and R²³ is 4-carboxybenzyl;

the compound of formula (IIa) wherein R²² is
20 5-chloro-2-thienyl and R²³ is 5-chloro-2-thienyl;

the compound of formula (IIa) wherein R²² is
2-quinolinylmethyl and R²³ is 2-quinolinylmethyl;

25 the compound of formula (IIa) wherein R²² is
2-propyl and R²³ is 2-picolinyl;

the compound of formula (IIa) wherein R²² is
3-benzyloxybenzyl and R²³ is 3-benzyloxybenzyl;

30

the compound of formula (IIa) wherein R²² is
4-phenylbenzyl and R²³ is phenylbenzyl;

the compound of formula (IIa) wherein R²² is
35 2-adamantylethyl and R²³ is 2-adamantylethyl;

the compound of formula (IIa) wherein R^{22} is hydrogen and R^{23} is cyclopropylmethyl;

5 the compound of formula (IIa) wherein R^{22} is 2-picolinyl and R^{23} is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is hydrogen;

10 the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is 2-picolinyl;

the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is 4-picolinyl;

15 the compound of formula (IIa) wherein R^{22} is 3-benzyloxybenzyl and R^{23} is 3-benzyloxybenzyl;

the compound of formula (IIa) wherein R^{22} is 3-cyclopropylmethoxybenzyl and R^{23} is 3-cyclopropylmethoxybenzyl;

20 the compound of formula (IIa) wherein R^{22} is 3-ethoxybenzyl and R^{23} is 3-ethoxybenzyl;

25 the compound of formula (IIa) wherein R^{22} is 4-benzyloxybenzyl and R^{23} is 4-benzyloxybenzyl.

the compound of formula (IIa) wherein R^{22} is 3-hydroxybenzyl and R^{23} is 3-hydroxybenzyl;

30 the compound of formula (IIa) wherein R^{22} is 4-hydroxybenzyl and R^{23} is 4-hydroxybenzyl;

35 the compound of formula (IIa) wherein R^{22} is 3-hydroxymethylbenzyl and R^{23} is 3-hydroxymethylbenzyl;

the compound of formula (IIa) wherein R²² is 4-hydroxymethylbenzyl and R²³ is 4-hydroxymethylbenzyl;

5 the compound of formula (IIa) wherein R²² is 3-aminobenzyl and R²³ is 3-aminobenzyl;

the compound of formula (IIa) wherein R²² is 3-carboxylbenzyl and R²³ is 3-carboxylbenzyl;

10 the compound of formula (IIa) wherein R²² is 3-formylbenzyl and R²³ is 3-formylbenzyl;

the compound of formula (IIa) wherein R²² is 3-cyanobenzyl and R²³ is 3-cyanobenzyl;

15

the compound of formula (IIa) wherein R²² is 2-naphthylmethyl and R²³ is 2-naphthylmethyl;

20 the compound of formula (IIa) wherein R²² is n-butyl and R²³ is benzyl;

the compound of formula (IIa) wherein R²² is allyl and R²³ is cyclopropylmethyl;

25

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is cyclopropylmethyl

the compound of formula (IIa) wherein R²² is 3-methallyl and R²³ is benzyl;

30

the compound of formula (IIa) wherein R²² is benzyl and R²³ is ethyl;

35 the compound of formula (IIa) wherein R²² is benzyl and R²³ is 4-picolinyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 4-picolinyl;

5 the compound of formula (IIa) wherein R²² is benzyl and R²³ is cyclopentylmethyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cyclopentylmethyl;
10

the compound of formula (IIa) wherein R²² is benzyl and R²³ is n-propyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cinnamyl;
15

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 2-naphthylmethyl;

20 the compound of formula (IIa) wherein R²² is cyclopentylmethyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is benzyl and R²³ is 2-naphthylmethyl;
25

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 2-picolinyl;

the compound of formula (IIa) wherein R²² is 3-cyanobenzyl and R²³ is 3-cyanobenzyl;
30

the compound of formula (IIa) wherein R²² is 3-allyl and R²³ is 2-naphthylmethyl;

35 the compound of formula (IIa) wherein R²² is n-propyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is n-butyl and R^{23} is 2-naphthylmethyl;

5 the compound of formula (IIa) wherein R^{22} is H and R^{23} is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is 4-picolinyl and R^{23} is 2-naphthylmethyl;

10

the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is cyclopentylmethyl;

15 the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is 2-quinolinylmethyl;

the compound of formula (IIa) wherein R^{22} is 3-picolinyl and R^{23} is cyclopropylmethyl;

20 the compound of formula (IIa) wherein R^{22} is 3-picolinyl and R^{23} is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is 3-allyloxybenzyl and R^{23} is 3-allyloxybenzyl;

25

the compound of formula (IIa) wherein R^{22} is 3-allyloxybenzyl and R^{23} is 3-hydroxybenzyl;

30 the compound of formula (IIa) wherein R^{22} is 3-picolinyl and R^{23} is 3-picolinyl;

the compound of formula (IIa) wherein R^{22} is 2-naphthylmethyl and R^{23} is 4-fluorobenzyl;

35 the compound of formula (IIa) wherein R^{22} is 3-carbomethoxybenzyl and R^{23} is 3-carbomethoxybenzyl;

the compound of formula (IIa) wherein R²² is 4-formylbenzyl and R²³ is 4-formylbenzyl;

5 the compound of formula (IIa) wherein R²² is 4-cyanobenzyl and R²³ is 4-cyanobenzyl;

the compound of formula (IIa) wherein R²² is 4-hydroxybenzyl and R²³ is n-propyl;

10

the compound of formula (IIa) wherein R²² is 3-hydroxybenzyl and R²³ is n-propyl;

the compound of formula (IIa) wherein R²² is 3-carboxybenzyl and R²³ is 3-carboxybenzyl;

15

the compound of formula (IIa) wherein R²² is cyclobutylmethyl and R²³ is cyclobutylmethyl;

20 the compound of formula (IIa) wherein R²² is cyclopentylmethyl and R²³ is cyclopentylmethyl;

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is 3-methylallyl;

25

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is cyclopentylmethyl;

the compound of formula (IIa) wherein R²² is 3-formaldoximebenzyl and R²³ is 3-formaldoximebenzyl;

30

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 3-hydroxybenzyl;

35 the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 4-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is 3-(N-methylamino)benzyl and R²³ is 3-(N-methylamino)benzyl;

5 the compound of formula (IIa) wherein R²² is 3-acetylbenzyl and R²³ is 3-acetylbenzyl;

the compound of formula (IIa) wherein R²² is 3-hydroxylaminobenzyl and R²³ is 3-hydroxylaminobenzyl;

10

the compound of formula (IIa) wherein R²² is 2-naphthylmethyl and R²³ is 3-hydroxybenzyl;

15 the compound of formula (IIa) wherein R²² is 4-hydroxymethylbenzyl and R²³ is 3-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is N-methyl-(3-amido)benzyl and R²³ is N-methyl-(3-amido)benzyl;

20

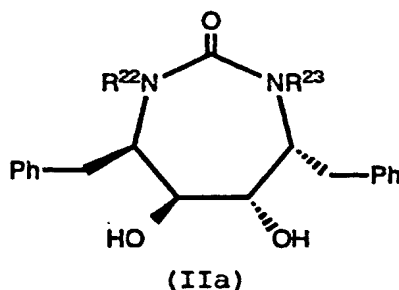
the compound of formula (IIa) wherein R²² is N-methyl-(3-amido)benzyl and R²³ is 3-(amidino)benzyl;

25 the compound of formula (IIa) wherein R²² is 3-(5-tetrazolyl)benzyl and R²³ is cyclopropylmethyl;

the compound of formula (IIa) wherein R²² is 3-(5-tetrazolyl)benzyl and R²³ is 3-(5-tetrazolyl) benzyl;

30 the compound of formula (IIa) wherein R²² is phenylmethyl-3-boronic acid and R²³ is phenylmethyl-3-boronic acid.

35 9. A compound of Claim 1 of formula (IIa):



which is selected from the group consisting of:

5

the compound of formula (IIa) wherein R²² is allyl and R²³ is allyl;

10 the compound of formula (IIa) wherein R²² is propyl and R²³ is propyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cyclopropylmethyl;

15 the compound of formula (IIa) wherein R²² is n-butyl and R²³ is n-butyl;

the compound of formula (IIa) wherein R²² is CH₂CH=CH(CH₃)₂ and R²³ is CH₂CH=CH(CH₃)₂;

20

the compound of formula (IIa) wherein R²² is i-pentyl and R²³ is i-pentyl;

25 the compound of formula (IIa) wherein R²² is 2-methylallyl and R²³ is 2-methylallyl;

the compound of formula (IIa) wherein R²² is n-pentyl and R²³ is n-pentyl;

30 the compound of formula (IIa) wherein R²² is benzyl and R²³ is benzyl;

the compound of formula (IIa) wherein R²² is allyl
and R²³ is isoprenyl;

5 the compound of formula (IIa) wherein R²² is
3-hydroxybenzyl and R²³ is 3-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is
4-hydroxybenzyl and R²³ is 4-hydroxybenzyl;

10

the compound of formula (IIa) wherein R²² is
3-hydroxymethylbenzyl and R²³ is 3-hydroxymethylbenzyl;

15 the compound of formula (IIa) wherein R²² is
4-hydroxymethylbenzyl and R²³ is 4-hydroxymethylbenzyl;

the compound of formula (IIa) wherein R²² is
3-aminobenzyl and R²³ is 3-aminobenzyl;

20 the compound of formula (IIa) wherein R²² is
3-carboxylbenzyl and R²³ is 3-carboxylbenzyl;

the compound of formula (IIa) wherein R²² is
3-formylbenzyl and R²³ is 3-formylbenzyl;

25

the compound of formula (IIa) wherein R²² is
3-cyanobenzyl and R²³ is 3-cyanobenzyl;

30 the compound of formula (IIa) wherein R²² is
2-naphthylmethyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is
n-butyl and R²³ is benzyl;

35 the compound of formula (IIa) wherein R²² is allyl
and R²³ is cyclopropylmethyl;

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is cyclopropylmethyl

5 the compound of formula (IIa) wherein R²² is 3-methallyl and R²³ is benzyl;

the compound of formula (IIa) wherein R²² is benzyl and R²³ is ethyl;

10

the compound of formula (IIa) wherein R²² is benzyl and R²³ is 4-picolinyl;

15 the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 4-picolinyl;

the compound of formula (IIa) wherein R²² is benzyl and R²³ is cyclopentylmethyl;

20 the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cyclopentylmethyl;

the compound of formula (IIa) wherein R²² is benzyl and R²³ is n-propyl;

25

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is cinnamyl;

30 the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R²² is cyclopentylmethyl and R²³ is 2-naphthylmethyl;

35 the compound of formula (IIa) wherein R²² is benzyl and R²³ is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is cyclopropylmethyl and R^{23} is 2-picolinyl;

5 the compound of formula (IIa) wherein R^{22} is 3-cyanobenzyl and R^{23} is 3-cyanobenzyl;

the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is 2-naphthylmethyl;

10

the compound of formula (IIa) wherein R^{22} is n-propyl and R^{23} is 2-naphthylmethyl;

15 the compound of formula (IIa) wherein R^{22} is n-butyl and R^{23} is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is H and R^{23} is 2-naphthylmethyl;

20 the compound of formula (IIa) wherein R^{22} is 4-picolinyl and R^{23} is 2-naphthylmethyl;

the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is cyclopentylmethyl;

25

the compound of formula (IIa) wherein R^{22} is 3-allyl and R^{23} is 2-quinolinylmethyl;

30 the compound of formula (IIa) wherein R^{22} is 3-picolinyl and R^{23} is cyclopropylmethyl;

the compound of formula (IIa) wherein R^{22} is 3-picolinyl and R^{23} is 2-naphthylmethyl;

35 the compound of formula (IIa) wherein R^{22} is 3-allyloxybenzyl and R^{23} is 3-allyloxybenzyl;

the compound of formula (IIa) wherein R^{22} is 3-allyloxybenzyl and R^{23} is 3-hydroxybenzyl;

5 the compound of formula (IIa) wherein R^{22} is 3-picolinyl and R^{23} is 3-picolinyl;

the compound of formula (IIa) wherein R^{22} is 2-naphthylmethyl and R^{23} is 4-fluorobenzyl;
10

the compound of formula (IIa) wherein R^{22} is 3-carbomethoxybenzyl and R^{23} is 3-carbomethoxybenzyl;

the compound of formula (IIa) wherein R^{22} is 4-formylbenzyl and R^{23} is 4-formylbenzyl;
15

the compound of formula (IIa) wherein R^{22} is 4-cyanobenzyl and R^{23} is 4-cyanobenzyl;

20 the compound of formula (IIa) wherein R^{22} is 4-formylbenzyl and R^{23} is 4-formylbenzyl;

the compound of formula (IIa) wherein R^{22} is 4-cyanobenzyl and R^{23} is 4-cyanobenzyl;
25

the compound of formula (IIa) wherein R^{22} is 4-formylbenzyl and R^{23} is 4-formylbenzyl;

the compound of formula (IIa) wherein R^{22} is 4-hydroxybenzyl and R^{23} is n-propyl;
30

the compound of formula (IIa) wherein R^{22} is 3-hydroxybenzyl and R^{23} is n-propyl;

35 the compound of formula (IIa) wherein R^{22} is 3-carboxybenzyl and R^{23} is 3-carboxybenzyl;

the compound of formula (IIa) wherein R²² is cyclobutylmethyl and R²³ is cyclobutylmethyl;

5 the compound of formula (IIa) wherein R²² is cyclopentylmethyl and R²³ is cyclopentylmethyl;

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is 3-methylallyl;

10

the compound of formula (IIa) wherein R²² is n-butyl and R²³ is cyclopentylmethyl;

15 the compound of formula (IIa) wherein R²² is 3-formaldoximebenzyl and R²³ is 3-formaldoximebenzyl;

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 3-hydroxybenzyl;

20

the compound of formula (IIa) wherein R²² is cyclopropylmethyl and R²³ is 4-hydroxybenzyl;

the compound of formula (IIa) wherein R²² is 3-(N-methylamino)benzyl and R²³ is 3-(N-methylamino)benzyl;

25

the compound of formula (IIa) wherein R²² is 3-acetylbenzyl and R²³ is 3-acetylbenzyl;

30 the compound of formula (IIa) wherein R²² is 3-hydroxylaminobenzyl and R²³ is 3-hydroxylaminobenzyl;

the compound of formula (IIa) wherein R²² is 2-naphthylmethyl and R²³ is 3-hydroxybenzyl;

35 the compound of formula (IIa) wherein R²² is 4-hydroxymethylbenzyl and R²³ is 3-hydroxybenzyl;

the compound of formula (IIa) wherein R^{22} is N-methyl-(3-amido)benzyl and R^{23} is N-methyl-(3-amido)benzyl;

5

the compound of formula (IIa) wherein R^{22} is N-methyl-(3-amido)benzyl and R^{23} is 3-(amidino)benzyl;

the compound of formula (IIa) wherein R^{22} is 3-(5-tetrazolyl)benzyl and R^{23} is cyclopropylmethyl;

10

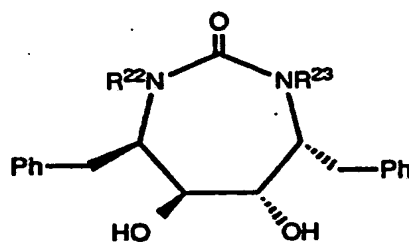
the compound of formula (IIa) wherein R^{22} is 3-(5-tetrazolyl)benzyl and R^{23} is 3-(5-tetrazolyl)benzyl;

15

the compound of formula (IIa) wherein R^{22} is phenylmethyl-3-boronic acid and R^{23} is phenylmethyl-3-boronic acid.

10. The compound of Claim 1 of formula (IIa):

20



25 which is selected from the group consisting of:

the compound of formula (IIa) wherein R^{22} is allyl and R^{23} is allyl;

30

the compound of formula (IIa) wherein R^{22} is cyclopropylmethyl and R^{23} is cyclopropylmethyl;

the compound of formula (IIa) wherein R^{22} is n-butyl and R^{23} is n-butyl;

5 the compound of formula (IIa) wherein R^{22} is $CH_2CH=CH(CH_3)_2$ and R^{23} is $CH_2CH=CH(CH_3)_2$;

the compound of formula (IIa) wherein R^{22} is propyl and R^{23} is propyl;

10

the compound of formula (IIa) wherein R^{22} is i-pentyl and R^{23} is i-pentyl;

15 the compound of formula (IIa) wherein R^{22} is benzyl and R^{23} is benzyl;

the compound of formula (IIa) wherein R^{22} is 3-hydroxybenzyl and R^{23} is 3-hydroxybenzyl;

20 the compound of formula (IIa) wherein R^{22} is 4-hydroxybenzyl and R^{23} is 4-hydroxybenzyl;

the compound of formula (IIa) wherein R^{22} is 3-hydroxymethylbenzyl and R^{23} is 3-hydroxymethylbenzyl;

25

the compound of formula (IIa) wherein R^{22} is 4-hydroxymethylbenzyl and R^{23} is 4-hydroxymethylbenzyl;

30 the compound of formula (IIa) wherein R^{22} is 3-aminobenzyl and R^{23} is 3-aminobenzyl;

the compound of formula (IIa) wherein R^{22} is 3-carboxybenzyl and R^{23} is 3-carboxybenzyl;

35 the compound of formula (IIa) wherein R^{22} is 3-formylbenzyl and R^{23} is 3-formylbenzyl

the compound of formula (IIa) wherein R^{22} is 3-formaldoximebenzyl and R^{23} is 3-formaldoximebenzyl;

5 the compound of formula (IIa) wherein R^{22} is 3-(N-methylamino)benzyl and R^{23} is 3-(N-methylamino)benzyl;

the compound of formula (IIa) wherein R^{22} is 3-acetylbenzyl and R^{23} is 3-acetylbenzyl;

10

the compound of formula (IIa) wherein R^{22} is 3-hydroxylaminobenzyl and R^{23} is 3-hydroxylaminobenzyl;

15 the compound of formula (IIa) wherein R^{22} is 2-naphthylmethyl and R^{23} is 3-hydroxybenzyl;

the compound of formula (IIa) wherein R^{22} is 4-hydroxymethylbenzyl and R^{23} is 3-hydroxybenzyl;

20 the compound of formula (IIa) wherein R^{22} is N-methyl-(3-amido)benzyl and R^{23} is N-methyl-(3-amido)benzyl;

25 the compound of formula (IIa) wherein R^{22} is N-methyl-(3-amido)benzyl and R^{23} is 3-(amidino)benzyl;

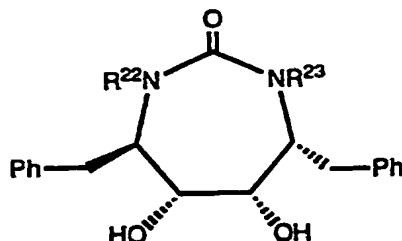
the compound of formula (IIa) wherein R^{22} is 3-(5-tetrazolyl)benzyl and R^{23} is cyclopropylmethyl;

30 the compound of formula (IIa) wherein R^{22} is 3-(5-tetrazolyl)benzyl and R^{23} is 3-(5-tetrazolyl)benzyl;

the compound of formula (IIa) wherein R^{22} is phenylmethyl-3-boronic acid and R^{23} is phenylmethyl-3-boronic acid.

35

11. A compound of Claim 1 of formula (IIb):

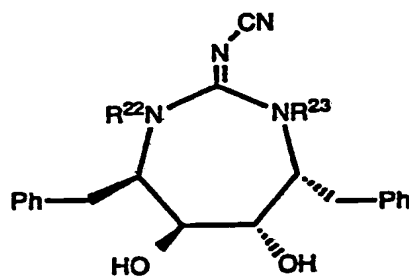


5 or a pharmaceutically acceptable salt or prodrug form thereof wherein:

R²² and R²³ are independently selected from the group consisting of: hydrogen, cyclopropylmethyl,
 10 CH₂(C₆H₄)-p-OCH₂C₆H₅, CH₂(C₆H₄)-p-OH, cyclopentylmethyl, allyl, n-butyl, beta-naphthylmethyl, benzyl, CH₂(C₆H₄)-m-OCH₂C₆H₅, p-nitrobenzyl, m-nitrobenzyl, CH₂(C₆H₄)-m-OH, CH₂(C₆H₄)-m-(CH₂OH), p-aminobenzyl, m-aminobenzyl,
 15 p-nitrilobenzyl, m-nitrilobenzyl, dimethylallyl, cyclohexylmethyl, cyclobutylmethyl, propyl, 3-methyl-1-butyl, carboxamidobenzyl, and formaldoximebenzyl.

20

12. A compound of Claim 1 of formula (Ib):



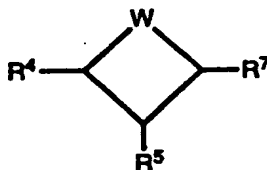
(Ib)

25

or a pharmaceutically acceptable salt or prodrug form thereof wherein:

5 R^{22} and R^{23} are independently selected from the group consisting of: hydrogen, cyclopropylmethyl, $CH_2(C_6H_4)-p-OCH_2C_6H_5$, $CH_2(C_6H_4)-p-OH$, cyclopentylmethyl, allyl, n-butyl, beta-naphthylmethyl, benzyl, $CH_2(C_6H_4)-m-OCH_2C_6H_5$,
 10 p-nitrobenzyl, m-nitrobenzyl, $CH_2(C_6H_4)-m-OH$, p-aminobenzyl, m-aminobenzyl, p-nitrilobenzyl, m-nitrilobenzyl, dimethylallyl, cyclohexylmethyl, cyclobutylmethyl, propyl, 3-methyl-1-butyl, carboxamidobenzyl, and formaldoximebenzyl.

15 13. A compound of Claim 1 of formula:



20 or a pharmaceutically acceptable salt or prodrug form thereof wherein:

R^4 and R^7 are independently selected from the following groups:

25

hydrogen;
 C₁-C₄ alkyl substituted with 0-3 R^{11} ;
 C₃-C₄ alkenyl substituted with 0-3 R^{11} ;

30 R^5 is $-OR^{20}$;

R^{11} is selected from one or more of the following:

keto, halogen, $-\text{CH}_2\text{NR}^{13}\text{R}^{14}$, $-\text{NR}^{13}\text{R}^{14}$, $-\text{OR}^{13}$, C₂-C₄ alkoxyalkyl, C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆ cycloalkyl;

5 aryl substituted with 0-3 R¹²; or

a heterocyclic ring system substituted with 0-2 R¹², composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

10

R¹², when a substituent on carbon, is selected from one or more of the following:

15 phenyl, benzyl, phenethyl, phenoxy, benzyloxy, halogen, C₁-C₄ alkyl, C₇-C₁₀ arylalkyl, C₁-C₄ alkoxy, $-\text{CO}_2\text{H}$, hydroxamic acid, hydrazide, oxime, boronic acid, sulfonamide, formyl, C₃-C₆ cycloalkoxy, $-\text{OR}^{13}$, C₁-C₄ alkyl substituted with $-\text{NR}^{13}\text{R}^{14}$, $-\text{NR}^{13}\text{R}^{14}$,
20 methylenedioxy, C₁-C₄ haloalkyl, C₁-C₄ alkylcarbonyl, C₁-C₄ alkylcarbonylamino, hydroxy, hydroxymethyl; or

a 5- or 6-membered heterocyclic ring containing from 1 to 4 heteroatoms selected from oxygen, nitrogen or
25 sulfur;

R¹², when a substituent on nitrogen, is selected from benzyl or methyl;

30 R¹³ is H, C₁-C₂ alkyl, or C₃-C₆ alkoxyalkyl;

R¹⁴ is OH, H or C₁-C₂ alkyl;

R¹³ and R¹⁴ can alternatively join to form $-(\text{CH}_2)_4-$,
35 $-(\text{CH}_2)_5-$, $-\text{CH}_2\text{CH}_2\text{N}(\text{R}^{15})\text{CH}_2\text{CH}_2-$, or $-\text{CH}_2\text{CH}_2\text{OCH}_2\text{CH}_2-$;

W is selected from:

- 5 -N(R²²)C(=Z)N(R²³)-;
 -C(R²⁵)(R²⁶)C(=Z)C(R²⁷)(R²⁸)-;
 -N(R²²)C(=Z)C(R²⁷)(R²⁸)-;
 -C(R²⁵)(R²⁶)C(=Z)O-;

wherein:

10

Z is O, S, or N-CN;

R²² and R²³ are independently selected from the following:

15

hydrogen;
C₁-C₄ alkyl substituted with 0-3 R³¹;
C₃-C₄ alkenyl substituted with 0-3 R³¹;

20 R²⁵ and R²⁷ are independently selected from the following:

hydrogen;
C₁-C₄ alkyl substituted with 0-3 R³¹;
25 C₃-C₄ alkenyl substituted with 0-3 R³¹;

R²⁶ and R²⁸ are hydrogen;

R³¹ is selected from one or more of the following:

30

keto, halogen, -CH₂NR¹³R¹⁴, -NR¹³R¹⁴, -OR¹³, C₂-C₄
alkoxyalkyl, C₁-C₄ alkyl, C₂-C₄ alkenyl, C₃-C₆
cycloalkyl;

35

aryl substituted with 0-3 R³²; or

a heterocyclic ring system substituted with 0-2 R^{32} , composed of 5 to 10 atoms including at least one nitrogen, oxygen or sulfur atom;

- 5 R^{32} , when a substituent on carbon, is selected from one or more of the following:

phenyl, benzyl, phenethyl, phenoxy, benzyloxy,
halogen, C₁-C₄ alkyl, C₇-C₁₀ arylalkyl, C₁-C₄ alkoxy,
10 -CO₂H, hydroxamic acid, hydrazide, oxime, boronic
acid, sulfonamide, formyl, C₃-C₆ cycloalkoxy, -OR¹³,
C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴,
methylenedioxy, C₁-C₄ haloalkyl, C₁-C₄ alkylcarbonyl,
C₁-C₄ alkylcarbonylamino, hydroxy, hydroxymethyl,
15 -C(R¹⁴)=N(OR¹⁴); or

a 5- or 6-membered heterocyclic ring containing from
1 to 4 heteroatoms selected from oxygen, nitrogen or
sulfur;

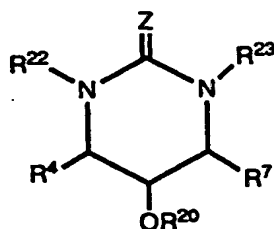
- 20 R^{32} , when a substituent on nitrogen, is selected from
benzyl or methyl;

provided that:

- 25 when R⁴ is hydrogen, R⁷ is not hydrogen;

when R⁴ is hydrogen, at least one of the following
is not hydrogen: R²², R²³, R²⁵, R²⁶, R²⁷ and R²⁸.

- 30 14. A compound of formula:



wherein:

R⁴ and R⁷ are independently selected from the following
5 groups:

hydrogen;

C₁-C₃ alkyl substituted with 0-1 R¹¹;

10

R²⁰ is hydrogen or any group that, when administered to a mammalian subject, cleaves to form a free hydroxyl;

R¹¹ is selected from one or more of the following:

15

halogen, -OR¹³, C₁-C₄ alkyl, C₃-C₅ cycloalkyl;

aryl substituted with 0-2 R¹²; or

20

a heterocyclic ring system chosen from pyridyl, pyrimidinyl, triazinyl, furanyl, thienyl, pyrrolyl, pyrazolyl, imidazolyl, tetrazolyl, benzofuranyl, indolyl, quinolinyl, isoquinolinyl;

25

R¹², when a substituent on carbon, is selected from one or more of the following:

benzyloxy, halogen, methyl, C₁-C₄ alkoxy, CF₃,

2-(1-morpholino)ethoxy, -CO₂H, hydroxamic acid,

30

hydrazide, oxime, cyano, boronic acid, sulfonamide,

formyl, C₃-C₆ cycloalkoxy, C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl; or

R¹², when a substituent on nitrogen, is methyl;

5

R¹³ is H or methyl;

R¹⁴ is OH, H or methyl;

10 R¹³ and R¹⁴ can alternatively join to form -(CH₂)₄-, -(CH₂)₅-, -CH₂CH₂N(R¹⁵)CH₂CH₂-, or -CH₂CH₂OCH₂CH₂-;

R²² and R²³ are independently selected from the following:

15

hydrogen;

C₁-C₄ alkyl substituted with 0-1 R³¹;

C₃-C₄ alkenyl substituted with 0-1 R³¹;

20 R³¹ is selected from one or more of the following:

halogen, -OR¹³, C₁-C₄ alkyl, C₃-C₅ cycloalkyl;

aryl substituted with 0-2 R³²; or

25

a heterocyclic ring system chosen from pyridyl, pyrimidinyl, triazinyl, furanyl, thienyl, pyrrolyl, pyrazolyl, imidazolyl, tetrazolyl, benzofuranyl, indolyl, quinolinyl, isoquinolinyl;

30

R³², when a substituent on carbon, is selected from one or more of the following:

35 benzyloxy, halogen, methyl, C₁-C₄ alkoxy, CF₃, 2-(1-morpholino)ethoxy, -CO₂H, hydroxamic acid, hydrazide, oxime, cyano, boronic acid, sulfonamide,

formyl, C₃-C₆ cycloalkoxy, C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl, -C(R¹⁴)=N(OR¹⁴); or

5 R³², when a substituent on nitrogen, is methyl;

provided that:

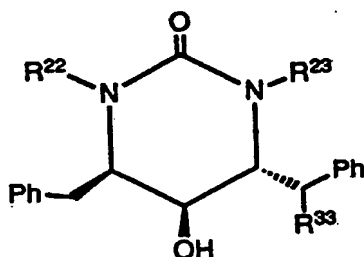
when R⁴ is hydrogen, R⁷ is not hydrogen;

10

when R⁴ is hydrogen, at least one of the following is not hydrogen: R²² and R²³.

15

15. A compound of Claim 1 of formula:



or a pharmaceutically acceptable salt or prodrug form thereof wherein:

20

R³³ is OH, halogen, H, N₃;

R²² and R²³ are independently selected from the group

25 consisting of:

hydrogen, allyl, propyl, cyclopropylmethyl, n-butyl, i-butyl, CH₂CH=CH(CH₃)₂, pyridylmethyl, methallyl, n-pentyl, i-pentyl, hexyl, benzyl, pyridylmethyl, isoprenyl, propargyl, picolinyl, methoxyethyl, cyclohexylmethyl, dimethyl-butyl, ethoxyethyl, methyl-

30

oxazolinylmethyl, naphthylmethyl,
methyloxazolinylmethyl, vinyloxyethyl,
pentafluorobenzyl, quinolinylmethyl, carboxybenzyl,
chloro-thienyl, picolinyl, benzyloxybenzyl,
5 phenylbenzyl, adamantylethyl, cyclopropylmethoxybenzyl,
ethoxybenzyl, hydroxybenzyl, hydroxymethylbenzyl,
aminobenzyl, formylbenzyl, cyanobenzyl, cinnamyl,
allyloxybenzyl, fluorobenzyl, cyclobutylmethyl,
formaldoximebenzyl, cyclopentylmethyl, nitrobenzyl,
10 nitrilobenzyl, carboxamidobenzyl, carbomethoxybenzyl,
and dimethylallyl.

16. A method for the treatment of viral infections
which comprises administering to a host in need of such
15 treatment a therapeutically effective amount of a
compound of Claim 1.

17. A method for the treatment of viral infections
which comprises administering to a host in need of such
20 treatment a therapeutically effective amount of a
compound of Claim 2.

18. A method for the treatment of viral infections
which comprises administering to a host in need of such
25 treatment a therapeutically effective amount of a
compound of Claim 3.

19. A method for the treatment of viral infections
which comprises administering to a host in need of such
30 treatment a therapeutically effective amount of a
compound of Claim 4.

20. A method for the treatment of viral infections
which comprises administering to a host in need of such
35 treatment a therapeutically effective amount of a
compound of Claim 5.

21. A method for the treatment of viral infections which comprises administering to a host in need of such treatment a therapeutically effective amount of a
5 compound of Claim 6.

22. A method for the treatment of viral infections which comprises administering to a host in need of such treatment a therapeutically effective amount of a
10 compound of Claim 7.

23. A method for the treatment of viral infections which comprises administering to a host in need of such treatment a therapeutically effective amount of a
15 compound of Claim 8.

24. A method for the treatment of viral infections which comprises administering to a host in need of such treatment a therapeutically effective amount of a
20 compound of Claim 9.

25. A method for the treatment of viral infections which comprises administering to a host in need of such treatment a therapeutically effective amount of a
25 compound of Claim 10.

26. A method for the treatment of viral infections which comprises administering to a host in need of such treatment a therapeutically effective amount of a
30 compound of Claim 11.

27. A method for the treatment of viral infections which comprises administering to a host in need of such treatment a therapeutically effective amount of a
35 compound of Claim 12.

28. A method for the treatment of viral infections which comprises administering to a host in need of such treatment a therapeutically effective amount of a compound of Claim 13.

5

29. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 1.

10

30. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 2.

15

31. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 3.

20

32. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 4.

25

33. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 5.

30

34. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 6.

35

35. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 7.

5

36. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 8.

10

37. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 9.

15

38. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 10.

20

39. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 11.

25

40. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 12.

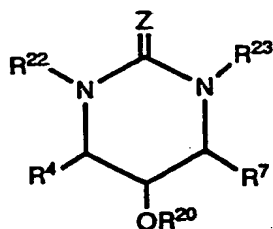
30

41. A pharmaceutical composition comprising a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound of Claim 13.

AMENDED CLAIMS

[received by the International Bureau
on 8 March 1993 (08.03.93);

original claims 14 and 15 amended; remaining claims unchanged (2 pages)]



wherein:

R⁴ and R⁷ are independently selected from the following
5 groups:

hydrogen;

C₁-C₃ alkyl substituted with 0-3 R¹¹;

10

R²⁰ is hydrogen or any group that, when administered to
a mammalian subject, cleaves to form a free hydroxyl;

R¹¹ is selected from one or more of the following:

15

halogen, -OR¹³, C₁-C₄ alkyl, C₃-C₅ cycloalkyl;

aryl substituted with 0-2 R¹²; or

20

a heterocyclic ring system chosen from pyridyl,
pyrimidinyl, triazinyl, furanyl, thienyl, pyrrolyl,
pyrazolyl, imidazolyl, tetrazolyl, benzofuranyl,
indolyl, quinolinyl, isoquinolinyl;

25

R¹², when a substituent on carbon, is selected from one
or more of the following:

benzyloxy, halogen, methyl, C₁-C₄ alkoxy, CF₃,
2-(1-morpholino)ethoxy, -CO₂H, hydroxamic acid,

30 hydrazide, oxime, cyano, boronic acid, sulfonamide,

formyl, C₃-C₆ cycloalkoxy, C₁-C₄ alkyl substituted with -NR¹³R¹⁴, -NR¹³R¹⁴, hydroxy, hydroxymethyl, -C(R¹⁴)=N(OR¹⁴); or

5 R³², when a substituent on nitrogen, is methyl;

provided that:

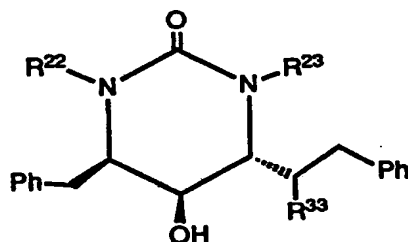
when R⁴ is hydrogen, R⁷ is not hydrogen;

10

when R⁴ is hydrogen, at least one of the following is not hydrogen: R²² and R²³.

15. A compound of Claim 1 of formula:

15



or a pharmaceutically acceptable salt or prodrug form thereof wherein:

20

R³³ is OH, halogen, H, N₃, or can alternatively be taken together with R²³ to form a direct bond;

R²² and R²³ are independently selected from the group
25 consisting of:

hydrogen, allyl, propyl, cyclopropylmethyl, n-butyl, i-butyl, CH₂CH=CH(CH₃)₂, pyridylmethyl, methallyl, n-pentyl, i-pentyl, hexyl, benzyl, pyridylmethyl,
30 isoprenyl, propargyl, picolinyl, methoxyethyl, cyclohexylmethyl, dimethyl-butyl, ethoxyethyl, methyl-

INTERNATIONAL SEARCH REPORT

PCT/US 92/08749

International Application No.

I. CLASSIFICATION OF SUBJECT MATTER (If several classification symbols apply, indicate all) ⁶		
According to International Patent Classification (IPC) or to both National Classification and IPC		
Int.Cl. 5 C07D243/04; C07D401/14;	C07D239/10; C07D413/14;	A61K31/55; C07D413/06; A61K31/505 C07D401/06
II. FIELDS SEARCHED		
Minimum Documentation Searched ⁷		
Classification System	Classification Symbols	
Int.Cl. 5	C07D	
Documentation Searched other than Minimum Documentation to the Extent that such Documents are Included in the Fields Searched ⁸		
III. DOCUMENTS CONSIDERED TO BE RELEVANT⁹		
Category ¹⁰	Citation of Document, ¹¹ with indication, where appropriate, of the relevant passages ¹²	Relevant to Claim No. ¹³
A	JOURNAL OF MEDICINAL CHEMISTRY vol. 33, no. 10, October 1990, WASHINGTON US pages 2687 - 2689 D.J. KEMPF ET AL. 'Structure-Based, C2 Symmetric Inhibitors of HIV Protease' see the whole document	1-41
<p>¹⁰ Special categories of cited documents: ¹⁰</p> <p>"A" document defining the general state of the art which is not considered to be of particular relevance</p> <p>"E" earlier document but published on or after the international filing date</p> <p>"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)</p> <p>"O" document referring to an oral disclosure, use, exhibition or other means</p> <p>"P" document published prior to the international filing date but later than the priority date claimed</p> <p>"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</p> <p>"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step</p> <p>"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</p> <p>"A" document member of the same patent family</p>		
IV. CERTIFICATION		
2	Date of the Actual Completion of the International Search 29 JANUARY 1993	Date of Mailing of this International Search Report - 8. 02. 93
International Searching Authority EUROPEAN PATENT OFFICE		Signature of Authorized Officer ALLARD M.S.

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 92/08749

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☐ Claims Nos.:
because they relate to subject matter not required to be searched by this Authority, namely:
Although claims 16-29 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compounds/compositions.
2. ☐ Claims Nos.:
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
Claims 1-3 and 13 encompass an enormous amount of compounds of dissimilar structures, for which no evidence is to be found in the description that they solve the problem underlying the application, namely the provision of compounds inhibiting retroviral protease. Pursuant to ✓
3. ☐ Claims Nos.:
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☐ No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/

Article 6 PCT and to the PCT Search Guidelines C-III, 2.1 and 3.7 these claims have only been searched insofar as they are sufficiently supported by the description.

Claims searched completely : 4-12,14,15,32-40
Claims searched incompletely : 1-3,13,29-31,41